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OM protein - protein search, using sw model

Run on: January 22, 2004, 16:31:15; Search time 8.44503 Seconds

(without alignments)

2650.131 Million cell updates/sec

Title:

US-09-830-972-32

Perfect score:

Sequence:

1 QASGEAGVSCLRENFAVYSV......ESEVAISEELVQKYSNSALG 141

Scoring table: BLOSUM62

Gapop 10.0 , Gapext 0.5

Searched:

1107863 seqs, 158726573 residues

Total number of hits satisfying chosen parameters:

1107863

Minimum DB seg length: 0

Maximum DB seq length: 2000000000

Post-processing: Minimum Match 0%

Maximum Match 100%

Listing first 45 summaries

Database :

A Geneseq 19Jun03:\*

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/SIDS1/qcqdata/geneseq/geneseqp-embl/AA1993.DAT:\* 14:

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/SIDS1/gcgdata/geneseq/geneseqp-emb1/AA1996.DAT:\* 17:

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/SIDS1/gcgdata/geneseq/geneseqp-emb1/AA2003.DAT:\*

Pred. No. is the number of results predicted by chance to have a score greater than or equal to the score of the result being printed, and is derived by analysis of the total score distribution.

# SUMMARIES

|        |       | ક     |        |    |          |                    |
|--------|-------|-------|--------|----|----------|--------------------|
| Result |       | Query |        |    |          |                    |
| No.    | Score | Match | Length | DB | ID       | Description        |
|        |       |       |        |    |          |                    |
| 1      | 680.5 | 96.5  | 522    | 21 | AAY71312 | Rat neurite growth |
| 2      | 510   | 72.3  | 199    | 23 | ABB81077 | Rat neurotransmitt |
| 3      | 503   | 71.3  | 118    | 23 | ABB89192 | Human polypeptide  |
| 4      | 503   | 71.3  | 199    | 19 | AAW53947 | Human NSPLP protei |
| 5      | 503   | 71.3  | 199    | 20 | AAY35903 | Extended human sec |
| 6      | 503   | 71.3  | 199    | 20 | AAW78313 | Fragment of human  |
| 7      | 503   | 71.3  | 199    | 21 | AAB12805 | Human NSPH protein |
| 8      | 503   | 71.3  | 199    | 22 | AAB82348 | Human NOGO-C prote |
| 9      | 503   | 71.3  | 199    | 23 | ABB81080 | Human neurotransmi |
| 10     | 503   | 71.3  | 199    | 23 | ABG30939 | Human NogoC protei |
| 11     | 499.5 | 70.9  | 199    | 21 | AAY71559 | Rat Nogo C/Nogo A  |
| 12     | 448   | 63.5  | 1178   | 21 | AAY71311 | Human neurite grow |
| 13     | 447   | 63.4  | 403    | 21 | AAY71563 | Rat Nogo A protein |
| 14     | 447   | 63.4  | 893    | 21 | AAY95012 | Human secreted pro |
| 15     | 447   | 63.4  | 983    | 24 | ABU11573 | Human MDDT polypep |
| 16     | 447   | 63.4  | 1162   | 21 | AAY71557 | Rat Nogo A truncat |
| 17     | 447   | 63.4  | 1163   | 21 | AAY71310 | Rat neurite growth |
| 18     | 447   | 63.4  | 1163   | 21 | AAY71384 | Alternative versio |
| 19     | 447   | 63.4  | 1163   | 23 | ABB81074 | Rat neurotransmitt |
| 20     | 447   | 63.4  | 1192   | 21 | AAY56967 | Human MAGI polypep |
| 21     | 447   | 63.4  | 1192   | 22 | AAU04591 | Human Nogo protein |
| 22     | 447   | 63.4  | 1192   | 22 | AAB82349 | Human NOGO-A prote |
| 23     | 447   | 63.4  | 1192   | 23 | ABP68600 | Human pancreatic c |
| 24     | 447   | 63.4  | 1192   | 23 | ABB81078 | Human neurotransmi |
| 25     | 447   | 63.4  | 1192   | 23 | ABG30938 | Human NogoA protei |
| 26     | 443   | 62.8  | 103    | 22 | AAE03980 | Human gene 42 enco |
| 27     | 443   | 62.8  | 200    | 22 | AAB64514 | Human secreted pro |
| 28     | 443   | 62.8  | 359    | 21 | AAY71558 | Rat Nogo A protein |
| 29     | 443   | 62.8  | 360    | 21 | AAY71383 | Rat neurite growth |
| 30     | 443   | 62.8  | 360    | 22 | AAE03987 | Human gene 42 enco |
| 31     | 443   | 62.8  | 360    | 23 | ABB81076 | Rat neurotransmitt |
| 32     | 443   | 62.8  | 361    | 21 | AAY71385 | Alternative versio |
| 33     | 443   | 62.8  | 373    | 21 | AAB24242 | Human Nogo B prote |
| 34     | 443   | 62.8  | 373    | 21 | AAY56969 | Human MAGI polypep |
| 35     | 443   | 62.8  | 373    | 21 | AAY53624 | A bone marrow secr |
| 36     | 443   | 62.8  | 373    | 22 | AAB82350 | Human NOGO-B prote |
| 37     | 443   | 62.8  | 373    | 23 | ABP68601 | Human pancreatic c |
| 38     | 443   | 62.8  | 373    | 23 | ABB81079 | Human neurotransmi |
| 39     | 443   | 62.8  | 373    | 23 | ABG30937 | Human NogoB protei |
| 40     | 443   | 62.8  | 373    | 23 | AAM47954 | Human RTN4B SEQ ID |
| 41     | 440   | 62.4  | 91     | 20 | AAY12360 | Human 5' EST secre |
| 42     | 439   | 62.3  | 291    | 22 | AAM93484 | Human polypeptide, |
| 43     | 410.5 | 58.2  | 642    | 19 | AAW58383 | Human secreted pro |
| 44     | 410.5 | 58.2  | 642    | 22 | AAB90682 | Human BG160_1 prot |
| 45     | 367   | 52.1  | 120    | 22 | AAE03939 | Human gene 42 enco |

```
RESULT 1
AAY71312
     AAY71312 standard; Protein; 522 AA.
ID
XX
AC
     AAY71312;
XX
DT
     02-NOV-2000
                  (first entry)
XX
DE
     Rat neurite growth inhibitor Nogo C.
XX
KW
     Rat; neurite growth inhibitor; Nogo C; neural cell; myelin; CNS;
     central nervous system; neoplastic disease; antiproliferative; glioma;
KW
     antisense gene therapy; neuroblastoma; menagioma; retinoblastoma;
KW
KW
     degenerative nerve disease; Alzheimer's disease; Parkinson's disease;
KW
     hyperproliferative disorder; benign dysproliferative disorder; diagnosis;
     psoriasis; tissue hypertrophy; neuronal regeneration; treatment;
KW
KW
     structural plasticity; screening.
XX
OS
     Rattus sp.
XX
FΗ
                     Location/Qualifiers
     Key
FT
                      1..39
     Region
FT
                      /note= "Sequence upstream to the N-terminus of
FT
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                      /label= Nogo C protein
FT
     Region
                      11..191
FT
                      /note= "Region specifically described in claim 16"
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     Region
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                      /note= "Sequence downstream to the C-terminus of
FT
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FT
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FT
                      /note= "C-terminal common region found in Nogo A, B
FT
                      and C isoforms "
FT
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FT
                      /note= "Encoded by TAG"
FT
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FT
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     Misc-difference 239
FT
FT
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     Misc-difference 371
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XX
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     WO200031235-A2.
XX
PD
     02-JUN-2000.
XX
PF
     05-NOV-1999;
                     99WO-US26160.
XX
PR
     06-NOV-1998;
                    98US-0107446.
XX
PΑ
     (SCHW/) SCHWAB M E.
     (CHEN/) CHEN M S.
PΑ
XX
PΙ
     Schwab ME,
                 Chen MS;
XX
DR
     WPI; 2000-400052/34.
DR
     N-PSDB; AAD01175.
XX
РΤ
     Nogo proteins and nucleic acids useful for treating neoplastic
PT
     disorders of the central nervous system and inducing regeneration of
PT
     neurons -
XX
PS
     Claim 7; Fig 14; 122pp; English.
XX
CC
     The present sequence is a rat Nogo C protein which is a
     potent neural cell growth inhibitor and is free of all central nervous
CC
CC
     system (CNS) myelin material with which it is natively associated.
CC
     Nogo proteins and fragments displaying neurite growth inhibitory
CC
     activity are used in the treatment of neoplastic disease of the CNS
CC
     e.g. glioma, glioblastoma, medulloblastoma, craniopharyngioma, ependyoma,
     pinealoma, haemangioblastoma, acoustic neuroma, oligodendroglioma,
CC
CC
     menagioma, neuroblastoma or retinoblastoma and degenerative nerve
CC
     diseases e.g. Alzheimer's and Parkinson's diseases. Therapeutics which
```

```
CC
     or beniqn dysproliferative disorders e.g. psoriasis and tissue
CC
     hypertrophy. Ribozymes or antisense Nogo nucleic acids can be used to
     inhibit production of Nogo protein to induce regeneration of neurons or
CC
     to promote structural plasticity of the CNS in disorders where neurite
CC
CC
     growth, regeneration or maintenance are deficient or desired.
CC
     The animal models can be used in diagnostic and screening methods for
CC
     predisposition to disorders and to screen for or test molecules which
CC
     can treat or prevent disorders or diseases of the CNS.
     Note: SEQ ID numbers 35-42 are referred in claim 32 and SEQ ID NO: 29
CC
CC
     in disclosure of the specification. However the specification does not
     include sequences for these SEQ ID numbers.
CC
XX
SO
     Sequence
               522 AA;
                         96.5%; Score 680.5; DB 21; Length 522;
  Query Match
                        98.6%; Pred. No. 1.3e-71;
  Best Local Similarity
  Matches 138; Conservative
                             0; Mismatches
                                               1; Indels
                                                              1; Gaps
QУ
           3 SGEAGVSCLRENFAVYSVSVGMHNL-LLLEGRSWQEMDGQKKHWKDKVVDLLYWRDIKKT 61
              Db
            4 SGEAGVSCLRENFAVYSVSVGMHNLXLLLEGRSWOEMDGOKKHWKDKVVDLLYWRDIKKT 63
QУ
           62 GVVFGASLFLLLSLTVFSIVSVTAYIALALLSVTISFRIYKGVIQAIAKSDEGHPFRAYL 121
              Db
           64 GVVFGASLFLLLSLTVFSIVSVTAYIALALLSVTISFRIYKGVIOAIOKSDEGHPFRAYL 123
Qу
          122 ESEVAISEELVOKYSNSALG 141
              Db
          124 ESEVAISEELVQKYSNSALG 143
RESULT 2
ABB81077
TD
     ABB81077 standard; Protein; 199 AA.
XX
· AC
     ABB81077;
XX
DT
     05-NOV-2002
                 (first entry)
XX
DE
     Rat neurotransmitter receptor protein Nogo-C.
XX
KW
     Nerve regeneration; neuroprotection; neuronal degeneration; CNS; PNS;
KW
     central nervous system; peripheral nervous system; tranquillizer; Nogo;
KW
     vulnerary; cerebroprotective; anti-tumour; antidiabetic; anticonvulsant;
KW
     nootropic; antiparkinsonian; ophthalmological; analgesic; hepatotropic;
KW
     osteopathic; vasotropic; nephrotropic; cytostatic; antigen; gene therapy;
KW
     neurotransmitter receptor; rat; receptor.
XX
OS
     Rattus norvegicus.
XX
PN
     US2002072493-A1.
XX
PD
     13-JUN-2002.
XX
PF
     28-JUN-2001; 2001US-0893348.
XX
```

promote Nogo activity can be used to treat or prevent hyperproliferative

```
PR
     19-MAY-1998;
                    98IL-0124500.
PR
     21-JUL-1998;
                    98WO-US14715.
     22-DEC-1998;
                    98US-0218277.
PR
     19-MAY-1999;
                    99US-0314161.
PR
XX
PA
     (YEDA ) YEDA RES & DEV CO LTD.
XX
PΙ
     Eisenbach-Schwartz M, Hauben E, Cohen IR, Beserman P, Mosonego A;
PΙ
     Moalem G;
XX
DR
     WPI; 2002-607255/65.
     N-PSDB; ABN86600.
DR
XX
PT
     Promoting nerve regeneration and preventing neuronal degeneration in
PT
     the central/peripheral nervous system from injury/disease, comprises
PT
     administering nervous system-specific activated T cells/antigen, or
PT
     analogs/peptides -
XX
PS
     Example 5; Page 48-49; 93pp; English.
XX
CC
     The invention relates to promoting nerve regeneration or conferring
CC
     neuroprotection and preventing or inhibiting neuronal degeneration in the
CC
     central/peripheral nervous system (NS). The method involves administering
CC
      NS-specific activated T cells, NS-specific antigen, its analogue or its
CC
     peptide, a nucleotide sequence the NS-specific antigen or its analogue or
     combinations. The method is useful for promoting nerve regeneration and
CC
CC
     preventing neuronal degeneration in central/peripheral nervous system
CC
     from injury/disease, where the injury is spinal cord injury, blunt
CC
     trauma, penetrating trauma, hemorrhagic stroke, ischaemic stroke or
CC
     damages caused by surgery such as tumour excision. The disease is not an
     autoimmune disease or neoplasm. The disease results in a degenerative
CC
CC
     process occurring in either gray or white matter or both. The disease
CC
     is diabetic neuropathy, senile dementia, Alzheimer's disease, Parkinson's
CC
     disease, facial nerve (Bell's) palsy, glaucoma, Huntington's chorea,
CC
     amyotrophic lateral sclerosis, non-arteritic optic neuropathy, and
CC
     vitamin deficiency, intervertebral disc herniation, prion diseases such
CC
     as Creutzfeldt-Jakob disease, carpal tunnel syndrome, peripheral
CC
     neuropathies associated with various diseases, including but not limited
CC
     to uremia, porphyria, hypoglycemia, Sjorgren Larsson syndrome, acute
CC
     sensory neuropathy, chronic ataxic neuropathy, biliary cirrhosis, primary
CC
     amyloidosis, obstructive lung diseases, acromegaly, malabsorption
CC
     syndromes, polycythemia vera, immunoglobulin (Ig) A- and IgG gamma-
CC
     pathies, complications of various drugs (e.g., metronidazole) and toxins
CC
     (e.g., alcohol or organophosphates), Charcot-Marie-Tooth disease, ataxia
CC
     telangectasia, Friedreich's ataxia, amyloid polyneuropathies,
CC
     adrenomyeloneuropathy, Giant axonal neuropathy, Refsum's disease, Fabry's
CC
     disease, or lipoproteinemia. The present sequence represents the rat
CC
     neurotransmitter receptor protein Nogo-C, an example of NS-specific
CC
     antigen.
· XX
SQ
     Sequence
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  Query Match
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  Best Local Similarity
                          99.0%;
                                  Pred. No. 5.3e-52;
  Matches 103; Conservative
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                                                                              0;
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Qу

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QУ
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RESULT 3
ABB89192
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XX
AC
    ABB89192;
XX
DT
    24-MAY-2002 (first entry)
XX
DE
    Human polypeptide SEQ ID NO 1568.
XX
KW
    Cytostatic; immunosuppressive; nootropic; neuroprotective; antiviral;
KW
     antiallergic; hepatotropic; antidiabetic; antiinflammatory; antiulcer;
    vulnerary; anticonvulsant; antibacterial; antifungal; antiparasitic;
KW
     cardiant; gene therapy; cancer; immune disorder; cardiovascular disorder;
KW
     neurological disease; infection; human; secreted protein.
KW
XX
OS
    Homo sapiens.
XX
PN
    WO200190304-A2.
XX
ÞD
    29-NOV-2001.
XX
PF
    18-MAY-2001; 2001WO-US16450.
XX
PR
    19-MAY-2000; 2000US-205515P.
XX
PΑ
     (HUMA-) HUMAN GENOME SCI INC.
XX
PI
    Birse CE, Rosen CA;
XX
DR
    WPI; 2002-122018/16.
DR
    N-PSDB; ABL89601.
XX
PT
    Novel 1405 isolated polypeptides, useful for diagnosis, treatment and
    prevention of neural, immune system, muscular, reproductive,
PT
    gastrointestinal, pulmonary, cardiovascular, renal and proliferative
PT
    disorders -
XX
PS
    Claim 11; SEQ ID NO 1568; 2081pp + Sequence Listing; English.
XX
CC
    The invention relates to novel genes (ABL89449-ABL90853) and proteins
CC
    (ABB89040-ABB90444) useful for preventing, treating or ameliorating
CC
    medical conditions e.g. by protein or gene therapy. The genes are
CC
    isolated from a range of human tissues disclosed in the specification.
CC
    The nucleic acids, proteins, antibodies and (ant)agonists are useful
CC
    in the diagnosis, treatment and prevention of: (a) cancer, e.g. breast
CC
    and ovarian cancer and other cancers of the adrenal gland, bone, bone
CC
    marrow, breast, gastrointestinal tract, liver, lung, or urogenital;
CC
     (b) immune disorders e.g. Addison's disease, allergies, autoimmune
```

```
disease, multiple sclerosis, rheumatoid arthritis and ulcerative
CC
CC
     colitis; (c) cardiovascular disorders such as myocardial ischaemias;
CC
     (d) wound healing; (e) neurological diseases e.g. cerebral anoxia and
CC
     epilepsy; and (f) infectious diseases such as viral, bacterial, fungal
CC
     and parasitic infections.
CC
    Note: The sequence data for this patent did not form part of the
    printed specification, but was obtained in electronic format directly
CC
CC
     from WIPO at ftp.wipo.int/pub/published pct sequences.
XX
SO
    Sequence
               118 AA;
  Query Match
                         71.3%; Score 503; DB 23; Length 118;
  Best Local Similarity
                        98.1%; Pred. No. 1.8e-51;
 Matches 102; Conservative
                               1; Mismatches
                                                1; Indels
                                                              0; Gaps 0;
Qу
          38 MDGQKKHWKDKVVDLLYWRDIKKTGVVFGASLFLLLSLTVFSIVSVTAYIALALLSVTIS 97
             Db
           1 MDGQKKNWKDKVVDLLYWRDIKKTGVVFGASLFLLLSLTVFSIVSVTAYIALALLSVTIS 60
          98 FRIYKGVIQAIAKSDEGHPFRAYLESEVAISEELVQKYSNSALG 141
QУ
             Db
          61 FRIYKGVIQAIQKSDEGHPFRAYLESEVAISEELVOKYSNSALG 104
RESULT 4
AAW53947
ID
    AAW53947 standard; Protein; 199 AA.
XX
AC
    AAW53947;
XX
DT
    24-JUL-1998 (first entry)
XX
DΕ
    Human NSPLP protein A.
XX
KW
    NSPLP; neuroendocrine-specific protein-like protein; human; gene therapy;
    neurodegenerative disease; amyotrophic lateral sclerosis; cancer.
KW
XX
OS
    Homo sapiens.
XX
PN
    WO9806841-A2.
XX
PD
    19-FEB-1998.
XX
PF
    24-JUL-1997;
                  97WO-US13469.
XX
PR
    12-AUG-1996;
                  96US-0700607.
XX
PΑ
    (INCY-) INCYTE PHARM INC.
XX
PΙ
    Au-Young J, Bandman O, Goli SK, Hillman J;
XX
DR
    WPI; 1998-159533/14.
DR
    N-PSDB; AAV23695.
XX
PT
    Human neuro-endocrine-specific protein-like proteins - useful for
PT
    diagnosis, monitoring and treatment of cancer and neuro-degenerative
```

haemolytic anaemia, autoimmune thyroiditis, diabetes mellitus, Crohn's

```
XX
    Claim 1; Page 38; 73pp; English.
PS
XX
    This sequence is a human neuroendocrine-specific protein-like
CC
CC
    protein (NSPLP) of the invention. Recombinant cells transformed with the
    DNA are used to express the NSPLP proteins, which are used to treat
CC
CC
    cancer and neurodegenerative diseases such as amyotrophic lateral
    sclerosis. Also antisense nucleic acids and antagonists of NSPLP can be
CC
CC
    used to inhibit activity of the NSPLP proteins. Antibodies specific for
CC
    NSPLP are used for diagnosis and monitoring treatment of diseases
    associated with NSPLP expression, in usual immunoassays, and to isolate
CC
CC
    NSPLP from natural sources. The NSPLP proteins, or their fragments can
CC
    also be used in drug screening to identify NSPLP antagonists. The nucleic
CC
    acid can be used diagnostically and for monitoring treatment (in
CC
    hybridisation or amplification assays); to isolate closely related
CC
    sequences; in gene therapy for both sense and antisense applications
CC
    (including use of ribozymes) and for mapping the natural genomic
CC
    sequence.
XX
SO
    Sequence
               199 AA;
 Query Match
                         71.3%; Score 503; DB 19; Length 199;
 Best Local Similarity
                         98.1%; Pred. No. 3.6e-51;
                                                1;
 Matches 102; Conservative
                             1; Mismatches
                                                    Indels
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                                                                          0;
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Qу
          98 FRIYKGVIQAIAKSDEGHPFRAYLESEVAISEELVOKYSNSALG 141
             Db
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XX
AC
    AAY35903;
XX
DT
    13-SEP-1999
                 (first entry)
XX
DE
    Extended human secreted protein sequence, SEQ ID NO. 152.
XX
KW
    Secreted protein; human; cytokine; cellular proliferation; cell movement;
KW
    cellular differentiation; immune system regulator; anti-inflammatory;
KW
    haematopoiesis regulator; tissue growth regulator; tumour inhibitor;
KW
    reproductive hormone regulator; chemotaxis; chemokinesis; gene therapy;
KW
    genetic disease.
XX
OS
    Homo sapiens.
XX
PN
    WO9931236-A2.
XX
PD
    24-JUN-1999.
XX
```

PT

disease

```
PF
     17-DEC-1998;
                   98WO-IB02122.
XX
PR
     10-AUG-1998;
                   98US-0096116.
     17-DEC-1997;
                   97US-0069957.
PR
PR
     09-FEB-1998;
                   98US-0074121.
     13-APR-1998:
                   98US-0081563.
PR
XX
     (GEST ) GENSET.
PA
XX
PΙ
     Bouqueleret L, Duclert A, Dumas Milne Edwards J;
XX
DR
    WPI; 1999-385906/32.
DR
     N-PSDB; AAX97587.
ХX
PT
    New isolated human secreted proteins
ХX
PS
     Claim 9; Page 185-186; 516pp; English.
XX
CC
    This sequence is encoded by an extended human secreted protein coding
CC
     sequence of the invention. The secreted proteins can be used in treating
CC
     or controlling a variety of human conditions. The secreted proteins may
CC
     act as cytokines or may affect cellular proliferation or differentiation
     or may act as immune system regulators, haematopoiesis regulators, tissue
CC
CC
    growth regulators, regulators of reproductive hormones or cell movement
CC
     or have chemotactic/chemokinetic, receptor/ligand, anti-inflammatory or
CC
     tumour inhibition activity. The DNAs can be used in forensic procedures
CC
     to identify individuals or in diagnostic procedures to identify
CC
     individuals having genetic diseases resulting from abnormal expression of
CC
     the genes corresponding to the extended cDNAs. They are also useful for
CC
     constructing a high resolution map of the human chromosomes. They can
CC
     also be used for gene therapy to control or treat genetic diseases.
XX
SQ
    Sequence
               199 AA;
 Query Match
                         71.3%; Score 503; DB 20; Length 199;
 Best Local Similarity
                         98.1%; Pred. No. 3.6e-51;
 Matches 102; Conservative
                              1; Mismatches
                                                1; Indels
                                                              0; Gaps
                                                                          0;
Qу
          38 MDGQKKHWKDKVVDLLYWRDIKKTGVVFGASLFLLLSLTVFSIVSVTAYIALALLSVTIS 97
             Db
           1 MDGQKKNWKDKVVDLLYWRDIKKTGVVFGASLFLLLSLTVFSIVSVTAYIALALLSVTIS 60
Qу
          98 FRIYKGVIQAIAKSDEGHPFRAYLESEVAISEELVOKYSNSALG 141
              Db
          61 FRIYKGVIQAIQKSDEGHPFRAYLESEVAISEELVOKYSNSALG 104
RESULT 6
AAW78313
ID
    AAW78313 standard; Protein; 199 AA.
XX
AC
    AAW78313;
XX
DT
    13-APR-1999 (first entry)
XX
DE
    Fragment of human secreted protein encoded by gene 69.
XX
```

```
KW
     Human; secreted protein; fusion protein; gene therapy; protein therapy;
     diagnosis; tissue; cancer; tumour; neurodegenerative disorder; leukaemia;
KW
     developmental abnormality; foetal deficiency; blood; allergy; renal;
KW
KW
     immune system; asthma; lymphocytic disease; brain; hepatic; lymphoma;
KW
     inflammation; ischaemic shock; Alzheimer's disease; restenosis; AIDS;
KW
     cognitive disorder; schizophrenia; prostate; obesity; osteoclast; thymus;
KW
     osteoporosis; arthritis; testis; lung; thyroiditis; thyroid; digestion;
KW
     endocrine; metabolism; regulation; malabsorption; gastritis; neoplasm.
XX
OS
     Homo sapiens.
XX
PN
     WO9856804-A1.
XX
PD
     17-DEC-1998.
XX
PF
     11-JUN-1998;
                     98WO-US12125.
XX
PR
                     97US-0061060.
     02-OCT-1997;
PR
     13-JUN-1997;
                     97US-0049547.
PR
     13-JUN-1997;
                     97US-0049548.
PR
     13-JUN-1997;
                     97US-0049549.
PR
     13-JUN-1997;
                     97US-0049550.
PR
     13-JUN-1997;
                     97US-0049606.
PR
     13-JUN-1997;
                     97US-0049607.
PR
     13-JUN-1997;
                     97US-0049608.
PR
     13-JUN-1997;
                     97US-0049609.
PR
     13-JUN-1997;
                     97US-0049610.
PR
     13-JUN-1997;
                     97US-0049611.
PR
     13-JUN-1997;
                     97US-0050566.
PR
     13-JUN-1997;
                     97US-0050901.
PR
     13-JUN-1997;
                     97US-0052989.
PR
     08-JUL-1997;
                     97US-0051919.
PR
     18-AUG-1997;
                     97US-0055984.
PR
     12-SEP-1997;
                     97US-0058665.
PR
     12-SEP-1997;
                     97US-0058668.
PR
     12-SEP-1997;
                     97US-0058669.
PR
     12-SEP-1997;
                     97US-0058750.
PR
     12-SEP-1997;
                     97US-0058971.
PR
     12-SEP-1997;
                     97US-0058972.
PR
     12-SEP-1997;
                     97US-0058975.
PR
     02-OCT-1997;
                     97US-0060834.
     02-OCT-1997;
PR
                     97US-0060841.
PR
     02-OCT-1997;
                     97US-0060844.
PR
     02-OCT-1997;
                     97US-0060865.
PR
     02-OCT-1997;
                     97US-0061059.
XX
PΑ
     (HUMA-) HUMAN GENOME SCI INC.
XX
PΙ
     Brewer LA, Ebner R, Ferrie AM, Feng P, Greene JM, Lafleur DW;
PΙ
     Moore PA,
                Ni J, Olsen HS, Rosen CA, Ruben SM, Shi Y, Young P;
PI
     Yu GL;
XX
DR
     WPI; 1999-080881/07.
DR
     N-PSDB; AAX04379.
XX
PT
     New isolated human genes and the secreted polypeptides they encode -
```

useful for diagnosis and treatment of e.g. cancers, neurological

PT

```
disorders, immune diseases, inflammation or blood disorders
XX
PS
    Disclosure; Page 62; 380pp; English.
XX
CC
    This sequence represents a fragment of a secreted human protein encoded
    by the nucleic acid molecule detailed in the descriptor line. The gene
CC
CC
    can be used to generate fusion proteins by linking to the gene to a
CC
    human immunoglobulin Fc portion (e.g. AAX04302) for increasing the
CC
    stability of the fused protein as compared to the human protein only.
CC
    The invention relates to 86 novel genes and their fragments (nucleic
CC
    acid sequences: AAX04311-X04410; amino acid sequences AAW78126-W78225)
CC
    which are useful for preventing, treating or ameliorating medical
CC
    conditions e.g. by protein or gene therapy. Also, pathological
CC
    conditions can be diagnosed by determining the amount of the new
    polypeptides in a sample or by determining the presence of mutations in
CC
    the new polynucleotides. Specific uses are described for each of the 86
CC
    polynucleotides, based on which tissues they are most highly expressed in
CC
CC
     (see AAX04311 for described uses).
XX
SQ
    Sequence
               199 AA;
 Query Match
                         71.3%; Score 503; DB 20; Length 199;
 Best Local Similarity
                        98.1%; Pred. No. 3.6e-51;
 Matches 102; Conservative 1; Mismatches
                                                1; Indels
                                                              0;
                                                                  Gaps
                                                                         0;
          38 MDGQKKHWKDKVVDLLYWRDIKKTGVVFGASLFLLLSLTVFSIVSVTAYIALALLSVTIS 97
QУ
             Db
           1 MDGQKKNWKDKVVDLLYWRDIKKTGVVFGASLFLLLSLTVFSIVSVTAYIALALLSVTIS 60
Qy
          98 FRIYKGVIQAIAKSDEGHPFRAYLESEVAISEELVOKYSNSALG 141
             Db
          61 FRIYKGVIQAIQKSDEGHPFRAYLESEVAISEELVQKYSNSALG 104
RESULT 7
AAB12805
ID
    AAB12805 standard; Protein; 199 AA.
XX
AC
    AAB12805;
XX
DT
    24-NOV-2000 (first entry)
XX
DE
    Human NSPH protein sequence SEQ ID NO:4.
XX
KW
    Human; neuroendocrine-specific protein; NSPH; NSPA; NSPB; NSPC.
XX
OS
    Homo sapiens.
XX
PN
    CN1253180-A.
XX
PD
    17-MAY-2000.
XX
PF
    30-OCT-1998;
                   98CN-0121473.
XX
PR
    30-OCT-1998;
                  98CN-0121473.
XX
PΑ
     (UYFU-) UNIV FUDAN.
```

PT

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XX
PΤ
     Yu L, Zhao Y, Zhang H;
XX
DR
     WPI; 2000-466537/41.
DR
     N-PSDB; AAA72981.
XX
     Specific protein of human neuroendocrine, coding sequence and its
PT
PT
     preparating process and application -
XX
PS
     Claim 4; Page 14-15; 21pp; Chinese.
XX
CC
     The present invention relates to a new member of the human
CC
     neuroendocrine specific protein family, designated NSPH. The present
CC
     sequence represents the human NSPH protein.
XX
               199 AA;
SO
     Sequence
  Query Match
                         71.3%; Score 503; DB 21; Length 199;
  Best Local Similarity
                         98.1%; Pred. No. 3.6e-51;
  Matches 102; Conservative
                              1; Mismatches
                                                1; Indels
                                                              0; Gaps
                                                                         0;
Qу
          38 MDGQKKHWKDKVVDLLYWRDIKKTGVVFGASLFLLLSLTVFSIVSVTAYIALALLSVTIS 97
             Db
           1 MDGQKKNWKDKVVDLLYWRDIKKTGVVFGASLFLLLSLTVFSIVSVTAYIALALLSVTIS 60
Qу
          98 FRIYKGVIQAIAKSDEGHPFRAYLESEVAISEELVOKYSNSALG 141
             Dh
          61 FRIYKGVIQAIQKSDEGHPFRAYLESEVAISEELVOKYSNSALG 104
RESULT 8
AAB82348
ID
    AAB82348 standard; Protein; 199 AA.
XX
АC
    AAB82348;
XX
    23-JUL-2001 (first entry)
DT
XX
DE
    Human NOGO-C protein.
XX
    NOGO-C; human; chromosome 2p21; neuropathy; spinal injury;
KW
    brain injury; stroke; neuronal degeneration; Alzheimer's disease;
KW
KW
    Parkinson's disease; neuromuscular disorder; psychiatric disorder;
    developmental disorder; neuroprotective; nootropic; neuroleptic;
KW
    antiparkinsonian; cerebroprotective; neuroleptic; diagnosis;
KW
KW
    therapy.
XX
OS
    Homo sapiens.
XX
PN
    WO200136631-A1.
XX
PD
    25-MAY-2001.
XX
PF
    14-NOV-2000; 2000WO-GB04345.
XX
PR
    15-NOV-1999;
                   99GB-0026995.
PR
    24-JAN-2000; 2000GB-0001550.
```

```
XX
PA
     (SMIK ) SMITHKLINE BEECHAM PLC.
XX
PΙ
     Michalovich D, Prinjha R;
XX
     WPI: 2001-343822/36.
DR
DR
     N-PSDB; AAF90323.
XX
PT
     New polypeptide designated NOGO-C is a splice variant of the human NOGO
РΤ
     gene and may be useful in the treatment of neural disorders including
РΤ
     Alzheimer's and Parkinson's diseases
XX
PS
     Claim 3; Page 25; 25pp; English.
XX
CC
     The present sequence is that of human NOGO-C, encoded by a novel
CC
     splice variant of the human NOGO gene on chromosome 2p21. 2 Other
CC
     splice variants, NOGO-A and NOGO-B, have previously been identified.
CC
     The invention provides NOGO-C polypeptides and polynucleotides, and
CC
     methods for producing such polypeptides by recombinant techniques.
     Also disclosed are methods for utilising NOGO-C polypeptides and
CC
CC
     polynucleotides in the treatment of diseases including neuropathies,
CC
     spinal injury, brain injury, stroke, neuronal degeneration, for
CC
     example Alzheimer's disease and Parkinson's disease, neuromuscular
CC
     disorders, psychiatric disorders and developmental disorders. Also
CC
     provided are methods for identifying agonists and agonists for
CC
     use in treating conditions associated with NOGO-C imbalance, and
CC
     diagnostic assays for detecting diseases associated with
CC
     inappropriate NOGO-C activity or levels.
XX
SO
     Sequence
               199 AA;
  Query Match
                         71.3%; Score 503; DB 22; Length 199;
 Best Local Similarity
                        98.1%; Pred. No. 3.6e-51;
 Matches 102; Conservative
                               1; Mismatches
                                               1; Indels
                                                              0; Gaps
                                                                          0;
Qу
          38 MDGQKKHWKDKVVDLLYWRDIKKTGVVFGASLFLLLSLTVFSIVSVTAYIALALLSVTIS 97
             Db
           1 MDGQKKNWKDKVVDLLYWRDIKKTGVVFGASLFLLLSLTVFSIVSVTAYIALALLSVTIS 60
Qу
          98 FRIYKGVIQAIAKSDEGHPFRAYLESEVAISEELVQKYSNSALG 141
             Db
          61 FRIYKGVIQAIQKSDEGHPFRAYLESEVAISEELVQKYSNSALG 104
RESULT 9
ABB81080
    ABB81080 standard; Protein; 199 AA.
XX
AC
    ABB81080;
XX
DT
    05-NOV-2002 (first entry)
XX
DE
    Human neurotransmitter receptor protein Nogo-C.
XX
KW
    Nerve regeneration; neuroprotection; neuronal degeneration; CNS; PNS;
KW
    central nervous system; peripheral nervous system; tranquillizer; Noqo;
KW
    vulnerary; cerebroprotective; anti-tumour; antidiabetic; anticonvulsant;
```

KW nootropic; antiparkinsonian; ophthalmological; analgesic; hepatotropic; KW osteopathic; vasotropic; nephrotropic; cytostatic; antigen; gene therapy; KW neurotransmitter receptor; human; receptor. XX OS Homo sapiens. XX PNUS2002072493-A1. XX PD 13-JUN-2002. XX PF 28-JUN-2001; 2001US-0893348. XX PR 19-MAY-1998; 98IL-0124500. 21-JUL-1998; 98WO-US14715. PR 22-DEC-1998; 98US-0218277. PR 19-MAY-1999; 99US-0314161. XX PA(YEDA ) YEDA RES & DEV CO LTD. ХХ PΙ Eisenbach-Schwartz M, Hauben E, Cohen IR, Beserman P, Mosonego A; PΙ Moalem G: XX DR WPI; 2002-607255/65. N-PSDB; ABN86601. DR XX PTPromoting nerve regeneration and preventing neuronal degeneration in PТ

Promoting nerve regeneration and preventing neuronal degeneration in the central/peripheral nervous system from injury/disease, comprises administering nervous system-specific activated T cells/antigen, or analogs/peptides -

Examples; Page 57-58; 93pp; English.

PΤ

PΤ

XX PS

XX CC

The invention relates to promoting nerve regeneration or conferring neuroprotection and preventing or inhibiting neuronal degeneration in the central/peripheral nervous system (NS). The method involves administering NS-specific activated T cells, NS-specific antigen, its analogue or its peptide, a nucleotide sequence the NS-specific antigen or its analogue or combinations. The method is useful for promoting nerve regeneration and preventing neuronal degeneration in central/peripheral nervous system from injury/disease, where the injury is spinal cord injury, blunt trauma, penetrating trauma, hemorrhagic stroke, ischaemic stroke or damages caused by surgery such as tumour excision. The disease is not an autoimmune disease or neoplasm. The disease results in a degenerative process occurring in either gray or white matter or both. The disease is diabetic neuropathy, senile dementia, Alzheimer's disease, Parkinson's disease, facial nerve (Bell's) palsy, glaucoma, Huntington's chorea, amyotrophic lateral sclerosis, non-arteritic optic neuropathy, and vitamin deficiency, intervertebral disc herniation, prion diseases such as Creutzfeldt-Jakob disease, carpal tunnel syndrome, peripheral neuropathies associated with various diseases, including but not limited to uremia, porphyria, hypoglycemia, Sjorgren Larsson syndrome, acute sensory neuropathy, chronic ataxic neuropathy, biliary cirrhosis, primary amyloidosis, obstructive lung diseases, acromegaly, malabsorption syndromes, polycythemia vera, immunoglobulin (Ig)A- and IgG gammapathies, complications of various drugs (e.g., metronidazole) and toxins (e.g., alcohol or organophosphates), Charcot-Marie-Tooth disease, ataxia telangectasia, Friedreich's ataxia, amyloid polyneuropathies,

```
disease, or lipoproteinemia. The present sequence represents the human
CC
CC
    neurotransmitter receptor protein Nogo-C, an example of NS-specific
CC
    antigen.
XX
SQ
    Sequence
               199 AA;
  Query Match
                        71.3%; Score 503; DB 23; Length 199;
                        98.1%; Pred. No. 3.6e-51;
  Best Local Similarity
  Matches 102; Conservative 1; Mismatches
                                                1; Indels
                                                             0; Gaps
                                                                         0;
Qу
          38 MDGQKKHWKDKVVDLLYWRDIKKTGVVFGASLFLLLSLTVFSIVSVTAYIALALLSVTIS 97
             Db
           1 MDGQKKNWKDKVVDLLYWRDIKKTGVVFGASLFLLLSLTVFSIVSVTAYIALALLSVTIS 60
          98 FRIYKGVIQAIAKSDEGHPFRAYLESEVAISEELVQKYSNSALG 141
QУ
             Db
          61 FRIYKGVIQAIQKSDEGHPFRAYLESEVAISEELVQKYSNSALG 104
RESULT 10
ABG30939
    ABG30939 standard; Protein; 199 AA.
XX
AC
    ABG30939;
XX
DT
    21-OCT-2002 (first entry)
XX
DE
    Human NogoC protein.
XX
KW
    Human; Nogo; BACE; acute neuronal injury; spinal injury; head injury;
KW
    stroke; peripheral nerve damage; neoplastic disorder; glioblastoma;
    neuroblastoma; hyperproliferative disorder; dysproliferative disorder;
KW
KW
    cirrhosis; psoriasis; keloid formation; fibrocystic condition; cancer;
KW
    tissue hypertrophy; central nervous system; axon regeneration; NogoC;
KW
    Nogo-associated disease; metastasis.
XX
OS
    Homo sapiens.
XX
    WO200257483-A2.
PN
XX
PD
    25-JUL-2002.
XX
    18-JAN-2002; 2002WO-GB00228.
PF
XX
PR
    18-JAN-2001; 2001GB-0001312.
XX
PA
     (GLAX ) GLAXO GROUP LTD.
PΑ
    (SMIK ) SMITHKLINE BEECHAM PLC.
XX
PΙ
    Blackstock WP, Hale RS, Prinjha R, Rowley A;
XX
DR
    WPI; 2002-599722/64.
DR
    N-PSDB; ABK90135.
XX
РΤ
    Identifying modulators of Nogo or BACE activity for treating acute
PT
    neuronal injuries, neoplastic or dysproliferative disorders, comprises
```

adrenomyeloneuropathy, Giant axonal neuropathy, Refsum's disease, Fabry's

```
PТ
     providing and monitoring interaction between Nogo and BACE polypeptides
PT
XX
PS
     Disclosure; Page 64; 68pp; English.
XX
CC
     The present invention relates to a new method of identifying modulators
     of Nogo function or BACE activity. The method involves providing Nogo and
CC
CC
     BACE polypeptides capable of binding with each other, monitoring the
CC
     interaction between these polypeptides, and determining if the test agent
CC
     is a modulator of Nogo or BACE activity. The method is useful in treating
CC
     acute neuronal injuries, such as spinal or head injury, stroke,
CC
     peripheral nerve damage, and in neoplastic (e.g. glioblastomas,
CC
     neuroblastomas), hyperproliferative or dysproliferative disorders (e.g.
CC
     cirrhosis, psoriasis, keloid formation, fibrocystic conditions, tissue
CC
     hypertrophy) of the central nervous system. The BACE polypeptide is
     useful in screening methods to identify agents that may act as modulators
CC
CC
     of BACE activity and in particular agents that may be useful in treating
CC
     Nogo-associated diseases. The modulators of Nogo or BACE polypeptides,
CC
     and the polynucleotide encoding the BACE polypeptide are useful in
CC
    manufacturing a medicament for the treatment or prevention of disorders
     responsive to the modulation of Nogo activity, in alleviating the
CC
CC
     symptoms or improving the condition of a patient suffering from this
CC
     disorder, in axon regeneration, or in preventing metastasis or spreading
CC
     of a cancer. The polynucleotide may also be an essential component in
CC
    assays, a probe, in recombinant protein synthesis, and in gene therapy
CC
     techniques. The present amino acid sequence represents the human NoqoC
CC
    protein of the invention.
XX
SO
    Sequence
              199 AA;
  Query Match
                         71.3%; Score 503; DB 23; Length 199;
  Best Local Similarity
                         98.1%; Pred. No. 3.6e-51;
 Matches 102; Conservative
                             1; Mismatches
                                               1; Indels
                                                                          0;
          38 MDGQKKHWKDKVVDLLYWRDIKKTGVVFGASLFLLLSLTVFSIVSVTAYIALALLSVTIS 97
Qу
             Db
           1 MDGQKKNWKDKVVDLLYWRDIKKTGVVFGASLFLLLSLTVFSIVSVTAYIALALLSVTIS 60
Qу
          98 FRIYKGVIQAIAKSDEGHPFRAYLESEVAISEELVQKYSNSALG 141
             Db
          61 FRIYKGVIQAIQKSDEGHPFRAYLESEVAISEELVQKYSNSALG 104
RESULT 11
AAY71559
ID
    AAY71559 standard; Protein; 199 AA.
XX
AC
    AAY71559;
XX
DT
    02-NOV-2000 (first entry)
XX
DE
    Rat Nogo C/Nogo A proteins derived fragment to construct mutant Nogo-C.
XX
KW
    Rat; neurite growth inhibitor; Nogo A; Nogo C; neural cell; myelin; CNS;
KW
    central nervous system; neoplastic disease; antiproliferative; glioma;
KW
    antisense gene therapy; neuroblastoma; menagioma; retinoblastoma;
KW
    degenerative nerve disease; Alzheimer's disease; Parkinson's disease;
```

KW hyperproliferative disorder; benign dysproliferative disorder; diagnosis; psoriasis; tissue hypertrophy; neuronal regeneration; treatment; KW structural plasticity; screening; mutant; mutein. KW XX OS Rattus sp. ХX FΗ Key Location/Qualifiers FTRegion 1..11 FT/note= "Corresponds to residues 40-50 of rat Nogo C FTprotein shown in AAY71312" FTRegion 12..199 FT/note= "Corresponds to residues 975-1162 of rat Nogo A FTprotein shown in AAY71310" XX PNWO200031235-A2. XX 02-JUN-2000. PD XX PF05-NOV-1999; 99WO-US26160. XXPR 06-NOV-1998; 98US-0107446. XXPΑ (SCHW/) SCHWAB M E. PΑ (CHEN/) CHEN M S. XX PΙ Schwab ME, Chen MS; XXDR WPI; 2000-400052/34. XX PTNogo proteins and nucleic acids useful for treating neoplastic PTdisorders of the central nervous system and inducing regeneration of PTneurons -XX PS Example; Page -; 122pp; English. XXCC The patent relates to neurite growth inhibitor Nogo which is free of CCall central nervous system (CNS) myelin material with which it is CCnatively associated. Nogo proteins and fragments displaying neurite CCgrowth inhibitory activity are used in the treatment of neoplastic CCdisease of the CNS e.g. glioma, glioblastoma, medulloblastoma, CC craniopharyngioma, ependyoma, pinealoma, haemangioblastoma, acoustic CC neuroma, oligodendroglioma, menagioma, neuroblastoma or retinoblastoma CCand degenerative nerve diseases e.g. Alzheimer's and Parkinson's CC diseases. Therapeutics which promote Nogo activity can be used to treat CCor prevent hyperproliferative or benign dysproliferative disorders e.q. CCpsoriasis and tissue hypertrophy. Ribozymes or antisense Nogo nucleic CCacids can be used to inhibit production of Nogo protein to induce CC regeneration of neurons or to promote structural plasticity of the CNS CC in disorders where neurite growth, regeneration or maintenance are deficient or desired. The animal models can be used in diagnostic and CC CCscreening methods for predisposition to disorders and to screen for or CC test molecules which can treat or prevent disorders or diseases of the CC CNS. The present sequence is derived by fusing two fragments from rat CC Nogo C and Nogo A proteins. The fragment is used in the construction of CC mutant Nogo-C which is composed of His-tag/T7-tag/Nogo-C N-terminus (11 aa) + Nogo-A sequence aa 975-1162. CC

Nogo A deletion mutants were used for mapping the inhibitory sites of

```
CC
     Nogo A sequence from amino acids 172-974, particularly amino acids
CC
     542-722. In addition, N-terminal region 1-171 was found to be inhibitory
CC
     to NIH 3T3 fibroblast spreading.
CC
     Note: The present sequence is not given in the specification but is
     derived from rat Nogo C sequence shown in AAY71312 and Nogo A sequence
CC
CC
     shown in AAY71310. SEQ ID numbers 35-42 are referred in claim 32 and
CC
     SEQ ID NO: 29 in disclosure of the specification. However, the
CC
     specification does not include sequences for these SEQ ID numbers.
XX
SO
     Sequence
               199 AA;
                         70.9%; Score 499.5; DB 21; Length 199;
  Query Match
  Best Local Similarity
                         98.1%; Pred. No. 9.3e-51;
  Matches 103; Conservative
                             0; Mismatches
                                                 1; Indels
                                                              1; Gaps
                                                                          1;
           38 MDGQKKHWKDK-VVDLLYWRDIKKTGVVFGASLFLLLSLTVFSIVSVTAYIALALLSVTI 96
Qу
              Db
           1 MDGQKKHWKDKSVVDLLYWRDIKKTGVVFGASLFLLLSLTVFSIVSVTAYIALALLSVTI 60
Qу
           97 SFRIYKGVIQAIAKSDEGHPFRAYLESEVAISEELVQKYSNSALG 141
              Db
           61 SFRIYKGVIQAIQKSDEGHPFRAYLESEVAISEELVOKYSNSALG 105
RESULT 12
AAY71311
     AAY71311 standard; Protein; 1178 AA.
XX
AC
     AAY71311;
XX
DT
     02-NOV-2000 (first entry)
XX
DE
     Human neurite growth inhibitor Nogo.
XX
KW
     Human; neurite growth inhibitor; Nogo; neural cell; myelin; CNS;
KW
     central nervous system; neoplastic disease; antiproliferative; glioma;
KW
     antisense gene therapy; neuroblastoma; menaqioma; retinoblastoma;
KW
     degenerative nerve disease; Alzheimer's disease; Parkinson's disease;
KW
     hyperproliferative disorder; beniqn dysproliferative disorder; diagnosis;
KW
     psoriasis; tissue hypertrophy; neuronal regeneration; treatment;
KW
     structural plasticity; screening.
XX
OS
     Homo sapiens.
XX
FH
                    Location/Qualifiers
FT
     Misc-difference 187
FT
                    /label= Unknown
FT
     Misc-difference 188
FT
                    /label= Unknown
    Misc-difference 189
FT
FT
                    /label= Unknown
FT
    Misc-difference 190
FT
                    /label= Unknown
FT
    Misc-difference 221
ĖΤ
                    /label= Unknown
FT
    Misc-difference 328
```

Nogo protein. Major inhibitory region was identified in the

```
FT
                     /label= Unknown
FT
     Misc-difference 477
FT
                     /label= Unknown
FT
                     994..1174
     Region
FT
                     /note= "Region specifically described in claim 16"
FT
     Region
FT
                     /note= "Region specifically described in claim 16"
FT
                     1079..1114
     Region
FT
                     /note= "Region specifically described in claim 16"
XX
PN
     WO200031235-A2.
ХX
PD
     02-JUN-2000.
XX
PF
     05-NOV-1999;
                    99WO-US26160.
XX
PR
     06-NOV-1998;
                    98US-0107446.
XX
PΑ
     (SCHW/) SCHWAB M E.
PΑ
     (CHEN/) CHEN M S.
XX
PΙ
     Schwab ME, Chen MS;
XX
DR
     WPI; 2000-400052/34.
XX
PT
     Nogo proteins and nucleic acids useful for treating neoplastic
PT
     disorders of the central nervous system and inducing regeneration of
PT
     neurons -
XX
PS
     Claim 11; Fig 13; 122pp; English.
XX
     The present sequence is a human Nogo protein which is a
CC
     potent neural cell growth inhibitor and is free of all central nervous
CC
CC
     system (CNS) myelin material with which it is natively associated. The
CC
     human Nogo sequence was derived by aligning human expressed sequence tags
CC
     (ESTs) e.g. AA158636, AA333267, AA081783, AA167765, AA322918, AA092565,
CC
     AA081525 and AA081840 with the rat Nogo sequence.
CC
     Nogo proteins and fragments displaying neurite growth inhibitory
CC
     activity are used in the treatment of neoplastic disease of the CNS
CC
     e.g. glioma, glioblastoma, medulloblastoma, craniopharyngioma, ependyoma,
CC
     pinealoma, haemangioblastoma, acoustic neuroma, oligodendroglioma,
CC
     menagioma, neuroblastoma or retinoblastoma and degenerative nerve
CC
     diseases e.g. Alzheimer's and Parkinson's diseases. Therapeutics which
     promote Nogo activity can be used to treat or prevent hyperproliferative
CC
CC
     or benign dysproliferative disorders e.g. psoriasis and tissue
CC
     hypertrophy. Ribozymes or antisense Nogo nucleic acids can be used to
CC
     inhibit production of Nogo protein to induce regeneration of neurons or
CC
     to promote structural plasticity of the CNS in disorders where neurite
CC
     growth, regeneration or maintenance are deficient or desired.
CC
     The animal models can be used in diagnostic and screening methods for
CC
     predisposition to disorders and to screen for or test molecules which
CC
     can treat or prevent disorders or diseases of the CNS.
CC
     Note: SEQ ID numbers 35-42 are referred in claim 32 and SEQ ID NO: 29
CC
     in disclosure of the specification. However the specification does not
CC
     include sequences for these SEQ ID numbers.
```

XX SQ

Sequence

1178 AA;

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  Best Local Similarity 74.6%; Pred. No. 1.3e-43;
  Matches
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                               3; Mismatches
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                                                            22; Gaps
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          12 RENFAVYSVSVGMHNLLLLEGRSWQEMDGQKKHWKDKVVDLLYWRDIKKTGVVFGASLFL 71
QУ
                 1::1 :1
                                             Db
         976 RSPSAIFSADLG-----
                                        ----KTSVVDLLYWRDIKKTGVVFGASLFL 1013
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QУ
             Db
        1014 LLSLTVFSIVSVTAYIALALLSVTISFRIYKGVIQAIQKSDEGHPFRAYLESEVAISEEL 1073
Qу
         132 VOKYSNSALG 141
             1074 VQKYSNSALG 1083
Db
RESULT 13
AAY71563
    AAY71563 standard; Protein; 403 AA.
XX
AC
    AAY71563;
XX
DT
    02-NOPR2000 (f
                       entry)
XX
DE
    Rat Nogo A protein fragment used in the construction of mutant EST.
XX
KW
    Rat; neurite growth inhibitor; Nogo A; neural cell; myelin; CNS;
KW
    central nervous system; neoplastic disease; antiproliferative; glioma;
KW
    antisense gene therapy; neuroblastoma; menagioma; retinoblastoma;
KW
    degenerative nerve disease; Alzheimer's disease; Parkinson's disease;
KW
    hyperproliferative disorder; benign dysproliferative disorder; diagnosis;
KW
    psoriasis; tissue hypertrophy; neuronal regeneration; treatment;
KW
    structural plasticity; screening; mutant; mutein.
XX
OS
    Rattus sp.
XX
PN
    WO200031235-A2.
XX
PD
    02-JUN-2000.
XX
PF
    05-NOV-1999;
                  99WO-US26160.
XX
PR
    06-NOV-1998;
                  98US-0107446.
XX
PA
    (SCHW/) SCHWAB M E.
PΑ
    (CHEN/) CHEN M S.
XX
PΙ
    Schwab ME, Chen MS;
XX
    WPI; 2000-400052/34.
DR
XX
PT
    Nogo proteins and nucleic acids useful for treating neoplastic
PT
    disorders of the central nervous system and inducing regeneration of
PT
    neurons -
XX
```

```
XX
CC
     The patent relates to neurite growth inhibitor Nogo which is free of
CC
     all central nervous system (CNS) myelin material with which it is
CC
     natively associated. Nogo proteins and fragments displaying neurite
CC
     growth inhibitory activity are used in the treatment of neoplastic
CC
     disease of the CNS e.g. glioma, glioblastoma, medulloblastoma,
CC
     craniopharyngioma, ependyoma, pinealoma, haemangioblastoma, acoustic
     neuroma, oligodendroglioma, menagioma, neuroblastoma or retinoblastoma
CC
     and degenerative nerve diseases e.g. Alzheimer's and Parkinson's
CC
     diseases. Therapeutics which promote Nogo activity can be used to treat
CC
CC
     or prevent hyperproliferative or benign dysproliferative disorders e.g.
CC
     psoriasis and tissue hypertrophy. Ribozymes or antisense Nogo nucleic
CC
     acids can be used to inhibh
                                      ction of Nogo protein180; induce
CC
     regeneration of neurons or to promote structural plasticity of the CNS
CC
     in disorders where neurite growth, regeneration or maintenance are
CC
     deficient or desired. The animal models can be used in diagnostic and
CC
     screening methods for predisposition to disorders and to screen for or
CC
     test molecules which can treat or prevent disorders or diseases of the
     CNS. The present sequence is a fragment of rat Nogo A protein shown in
CC
CC
     AAY71310, which is used in the construction of mutant EST. The mutant
CC
     is composed of His-tag/T7-tag/Nogo-A sequence aa 760-1162.
CC
     Nogo A deletion mutants were used for mapping the inhibitory sites of
CC
     Nogo protein. Major inhibitory region was identified in the
     Nogo A sequence from amino acids 172-974, particularly amino acids
CC
CC
     542-722. In addition, N-terminal region 1-171 was found to be inhibitory
CC
     to NIH 3T3 fibroblast spreading.
CC
    Note: The present sequence is not given in the specification but is
CC
     derived from rat Nogo A sequence shown in AAY71310. SEQ ID numbers 35-42
CC
     are referred in claim 32 and SEQ ID NO: 29 in disclosure of the
CC
     specification. However, the specification does not include sequences for
CC
     these SEQ ID numbers.
XX
SO
     Sequence
               403 AA;
  Query Match
                         63.4%;
                                 Score 447; DB 21; Length 403;
  Best Local Similarity
                         96.9%; Pred. No. 3.9e-44;
  Matches
           93; Conservative
                                0; Mismatches
                                                 3; Indels
                                                               0; Gaps
                                                                           0:
Qу
          46 KDKVVDLLYWRDIKKTGVVFGASLFLLLSLTVFSIVSVTAYIALALLSVTISFRIYKGVI 105
                Db
          214 KTSVVDLLYWRDIKKTGVVFGASLFLLLSLTVFSIVSVTAYIALALLSVTISFRIYKGVI 273
Qу
         106 QAIAKSDEGHPFRAYLESEVAISEELVOKYSNSALG 141
              Db
         274 QAIQKSDEGHPFRAYLESEVAISEELVQKYSNSALG 309
RESULT 14
    AAY95012 standard; Protein; 893 AA.
XX
AC
    AAY95012;
XX
    19-JUN-2000 (first entry)
DT
XX
DE
    Human secreted protein vb22 1, SEQ ID NO:64.
```

PS

Example; Page -; 122pp; English.

```
XX
ΚW
     Human; secreted protein; cancer; tumour; cardiovascular disorder;
KW
     blood disorder; haemophilia; autoimmune disease; diabetes; inflammation:
KW
     infection; fungal; bacterial; viral; HIV; allergy; arthritis;
     neurodegenerative disease; asthma; contraceptive.
KW
XX
OS
     Homo sapiens.
XX
     WO200011015-A1.
ΡN
XX
PD
     02-MAR-2000.
XX
PF
     24-AUG-1999;
                    99WO-US19351.
XX
PR
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     24-AUG-1998:
     24-AUG-1998;
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PR
     09-SEP-1998;
                    98US-0099618.
PR
     28-SEP-1998;
                    98US-0102092.
PR
     25-NOV-1998;
                    98US-0109978.
PR
     23-DEC-1998;
                    98US-0113645.
PR
     23-DEC-1998;
                    98US-0113646.
PR
     23-AUG-1999;
                    99US-0379246.
XX
РΔ
     (ALPH-) ALPHAGENE INC.
XX
     Valenzuela D, Yuan O, Hoffman H, Hall J, Rapiejko P;
ΡI
XX
DR
     WPI; 2000-224657/19.
XX
PT
     New secreted or transmembrane proteins and polynucleotides encoding
PT
     them, useful for treating neurodegenerative disorders, autoimmune
PT
     diseases and cancer -
XX
PS
     Claim 73; Page 322-325; 357pp; English.
ХX
CC
     The invention relates to 40 human secreted proteins (AAY94981-Y95020),
     and cDNA sequences encoding them (AAA23423-A23462). The secreted
CC
CC
     proteins of the invention include those that are thought to be only
CC
     partially secreted, i.e., transmembrane proteins. The proteins of the
CC
     invention may exhibit one or more activities selected from the following:
CC
     cytokine activity; cell proliferation; differentiation; immune
CC
     modulation; haematopoiesis regulation; tissue growth activity;
CC
     activin/inhibin activity; chemotactic/chemokinetic activity; haemostatic
CC
     and thrombolytic activity; anti-inflammatory activity; and tumour
CC
     inhibition activity. The proteins may be administered to patients as
CC
     vaccines, and the nucleotides may be used as part of a gene therapy
CC
     regime. Diseases or conditions that may be treated using the proteins or
CC
    nucleotides of the invention include autoimmune diseases; genetic
CC
    disorders; haemophilia; cardiovascular diseases; cancer; bacterial,
CC
     fungal and viral infections, especially HIV; multiple sclerosis;
CC
     rheumatoid arthritis; pulmonary inflammation; Guillain-Barre syndrome;
     insulin dependent diabetes mellitus; and allergic reactions such as
CC
```

asthma and anaemia. They may also be used for treating wounds, burns,

disease, Parkinson's disease, Huntington's disease and amyotrophic

lateral sclerosis (ALS). Proteins with activin/inhibin activity may

ulcers, osteoporosis, osteoarthritis, periodontal diseases, Alzheimer's

additionally be useful as contraceptives. Nucleic acid sequences of the

CC

CC

CC

CC

```
CC
     diagnostic primers and probes. The present sequence represents one of the
CC
     40 proteins of the invention.
XX
SQ
     Sequence
               893 AA;
  Query Match
                         63.4%; Score 447; DB 21; Length 893;
  Best Local Similarity
                         96.9%; Pred. No. 1.1e-43;
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Qу
                Db
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Qу
              Db
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XX
AC
     ABU11573;
XX
DT
     12-FEB-2003 (first entry)
XX
DE
     Human MDDT polypeptide SEQ ID 520.
XX
     MDDT; human; disease detection and treatment molecule polypeptide;
KW
KW
     anti-inflammatory; immunosuppressive; osteopathic; cytostatic; anti-HIV;
     haemostatic; nephrotropic; antianaemic; antipsoriatic; hepatotropic;
KW
     gene therapy; protein replacement therapy; cell proliferative disorder;
KW
KW
     cancer; adenocarcinoma; leukaemia; lymphoma; melanoma; myeloma; sarcoma;
     anaemia; Crohn's disease; acquired immunodeficiency syndrome; AIDS;
KW
KW
     Goodpasture's syndrome; inflammation; osteoporosis; thrombocytopaenia;
KW
    psoriasis; hepatitis.
XX
OS
    Homo sapiens.
XX
PN
    WO200279449-A2.
ХX
PD
    10-OCT-2002.
XX
PF
    27-MAR-2002; 2002WO-US09944.
XX
    28-MAR-2001; 2001US-279619P.
PR
PR
    29-MAR-2001; 2001US-280067P.
PR
    29-MAR-2001; 2001US-280068P.
PR
    16-MAY-2001; 2001US-291280P.
PR
    17-MAY-2001; 2001US-291829P.
PR
    17-MAY-2001; 2001US-291849P.
    19-JUN-2001; 2001US-299428P.
PR
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    20-JUN-2001; 2001US-299776P.
PR
    20-JUN-2001; 2001US-300001P.
XX
PΑ
     (INCY-) INCYTE GENOMICS INC.
```

invention may be used in chromosome mapping, and as a source of

```
XX
PΙ
     Daffo A, Jones AL, Tran AB, Dahl CR, Gietzen D,
     Dufour GE, Hillman JL, Yu JY, Tuason O, Yap PE, Amshey SR; Daugherty SC, Dam TC, Liu TF, Nguyen DA, Kleefeld Y, Gerstin EH;
PI
PΙ
     Peralta CH, David MH, Lewis SA, Chen AJ, Panzer SR, Harris B;
ΡI
ΡI
     Flores V, Marwaha R, Lo A, Lan RY, Urashka ME;
XX
DR
     WPI; 2003-058431/05.
DR
     N-PSDB; ABX34563.
XX
PT
     New purified disease detection and treatment molecule proteins and
     polynucleotides, useful for diagnosing, treating or preventing cancers
PT
PT
     (e.g. leukemia or sarcoma), anemia, Crohn's disease, AIDS, osteoporosis
PT
     or hepatitis -
XX
     Claim 27; SEQ ID NO 520; 339pp + Sequence Listing; English.
PS
XX
CC
     This invention describes a novel disease detection and treatment molecule
     polypeptide (MDDT) which has anti-inflammatory, immunosuppressive,
CC
CC
     osteopathic, cytostatic, anti-HIV, haemostatic, nephrotropic,
CC
     antianaemic, antipsoriatic and hepatotropic activity. The polynucleotides
CC
     and the polypeptides of the invention can be used for gene therapy,
CC
     protein replacement therapy and are useful for treating a variety of
CC
     diseases or conditions. These polypeptides or polynucleotides are
CC
     particularly useful for diagnosing, treating or preventing cell
CC
     proliferative disorders (e.g. cancers including adenocarcinoma,
CC
     leukaemia, lymphoma, melanoma, myeloma or sarcoma), anaemia, Crohn's
CC
     disease, acquired immunodeficiency syndrome (AIDS), Goodpasture's
     syndromes, inflammation, osteoporosis, thrombocytopaenia, psoriasis or
CC
     hepatitis. ABU11450-ABU11845 represent the MDDT polynucleotides encoded
CC
CC
     by ABU11450-ABU11845, described in the disclosure of the invention.
CC
     NOTE: The smqueapeedata for this p
                                              d not form part of the printed
CC
     specification, but was obtained in electronic format from WIPO at
CC
     ftp.wipo.int/pub/published pct sequences.
ХX
SO
     Sequence
               983 AA;
  Query Match
                         63.4%; Score 447; DB 24; Length 983;
  Best Local Similarity
                         96.9%; Pred. No. 1.3e-43;
  Matches
          93; Conservative
                              0; Mismatches
                                                 3; Indels
                                                                0;
                                                                   Gaps
Qу
          46 KDKVVDLLYWRDIKKTGVVFGASLFLLLSLTVFSIVSVTAYIALALLSVTISFRIYKGVI 105
                Db
         793 KTSVVDLLYWRDIKKTGVVFGASLFLLLSLTVFSIVSVTAYIALALLSVTISFRIYKGVI 852
Qу
         106 QAIAKSDEGHPFRAYLESEVAISEELVQKYSNSALG 141
              853 QAIQKSDEGHPFRAYLESEVAISEELVQKYSNSALG 888
Db
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Search completed: January 22, 2004, 16:36:41 Job time: 13.445 secs

### GenCore version 5.1.6 Copyright (c) 1993 - 2004 Compugen Ltd.

OM protein - protein search, using sw model

Run on: January 22, 2004, 16:31:15; Search time 3.42077 Seconds

(without alignments)

1744.001 Million cell updates/sec

Title:

US-09-830-972-32

Perfect score: 705

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Scoring table: BLOSUM62

Gapop 10.0 , Gapext 0.5

Searched:

328717 seqs, 42310858 residues

Total number of hits satisfying chosen parameters: 328717

Minimum DB seq length: 0

Maximum DB seq length: 2000000000

Post-processing: Minimum Match 0%

Maximum Match 100%

Listing first 45 summaries

Database :

Issued Patents AA:\*

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2: /cgn2\_6/ptodata/1/iaa/5B\_COMB.pep:\*

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4: /cgn2\_6/ptodata/1/iaa/6B COMB.pep:\*

5: /cgn2 6/ptodata/1/iaa/PCTUS\_COMB.pep:\*

6: /cgn2\_6/ptodata/1/iaa/backfiles1.pep:\*

Pred. No. is the number of results predicted by chance to have a score greater than or equal to the score of the result being printed, and is derived by analysis of the total score distribution.

### SUMMARIES

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| Result |       | Query |        |    |                    |                   |
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| 1      | 503   | 71.3  | 199    | 2  | US-08-700-607-1    | Sequence 1, Appli |
| 2      | 349   | 49.5  | 208    | 2  | US-08-700-607-7    | Sequence 7, Appli |
| 3      | 348   | 49.4  | 267    | 2  | US-08-700-607-8    | Sequence 8, Appli |
| 4      | 337   | 47.8  | 356    | 2  | US-08-700-607-6    | Sequence 6, Appli |
| 5      | 337   | 47.8  | 776    | 2  | US-08-700-607-5    | Sequence 5, Appli |
| 6      | 305   | 43.3  | 241    | 2  | US-08-700-607-3    | Sequence 3, Appli |
| 7      | 227   | 32.2  | 168    | 4  | US-09-149-476-563  | Sequence 563, App |
| 8      | 99    | 14.0  | 80     | 3  | US-08-905-223-411  | Sequence 411, App |
| 9      | 75    | 10.6  | 593    | 4  | US-09-328-352-4866 | Sequence 4866, Ap |
| 10     | 72.5  | 10.3  | 598    | 2  | US-08-853-659A-53  | Sequence 53, Appl |
| 11     | 71.5  | 10.1  | 154    | 1  | US-08-366-783-5    | Sequence 5, Appli |

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| 13 | 70   | 9.9 | 563  | 4 | US-09-422-936-79         | Sequence 79, Appl |
| 14 | 70   | 9.9 | 619  | 3 | US-08-262-220-6          | Sequence 6, Appli |
| 15 | 70   | 9.9 | 619  | 3 | US-08-471-733-6          | Sequence 6, Appli |
| 16 | 70   | 9.9 | 619  | 3 | US-08-468-878-6          | Sequence 6, Appli |
| 17 | 70   | 9.9 | 619  | 3 | US-08-750-494-6          | Sequence 6, Appli |
| 18 | 70   | 9.9 | 619  | 4 | US-08-470-638-6          | Sequence 6, Appli |
| 19 | 70   | 9.9 | 844  | 4 | US-09-422-936-47         | Sequence 47, Appl |
| 20 | 70   | 9.9 | 844  | 4 | US-09-422-936-51         | Sequence 51, Appl |
| 21 | 70   | 9.9 | 886  | 4 | US-09-422-936-77         | Sequence 77, Appl |
| 22 | 70   | 9.9 | 892  | 4 | US-09-422-936-75         | Sequence 75, Appl |
| 23 | 70   | 9.9 | 899  | 4 | US-09-422-936-71         | Sequence 71, Appl |
| 24 | 70   | 9.9 | 960  | 4 | US-09-422-936-45         | Sequence 45, Appl |
| 25 | 70   | 9.9 | 961  | 4 | US-09-422-936-49         | Sequence 49, Appl |
| 26 | 70   | 9.9 | 961  | 4 | US-09-914-259-14         | Sequence 14, Appl |
| 27 | 69   | 9.8 | 621  | 3 | US-08-262-220-8          | Sequence 8, Appli |
| 28 | 69   | 9.8 | 621  | 3 | US-08-471-733-8          | Sequence 8, Appli |
| 29 | 69   | 9.8 | 621  | 3 | US-08-468 <b>-</b> 878-8 | Sequence 8, Appli |
| 30 | 69   | 9.8 | 621  | 3 | US-08-750-494-8          | Sequence 8, Appli |
| 31 | 69   | 9.8 | 621  | 4 | US-08-470-638-8          | Sequence 8, Appli |
| 32 | 68   | 9.6 | 344  | 4 | US-09-107-532A-6886      | Sequence 6886, Ap |
| 33 | 68   | 9.6 | 1447 | 3 | US-09-041-886-25         | Sequence 25, Appl |
| 34 | 68   | 9.6 | 1447 | 5 | PCT-US94-05277-2         | Sequence 2, Appli |
| 35 | 67.5 | 9.6 | 659  | 4 | US-09-328-352-6021       | Sequence 6021, Ap |
| 36 | 67   | 9.5 | 231  | 4 | US-09-198-452A-419       | Sequence 419, App |
| 37 | 67   | 9.5 | 507  | 4 | US-09-328-352-7742       | Sequence 7742, Ap |
| 38 | 66.5 | 9.4 | 249  | 4 | US-09-107-532A-6706      | Sequence 6706, Ap |
| 39 | 66   | 9.4 | 445  | 4 | US-09-328-352-4714       | Sequence 4714, Ap |
| 40 | 65.5 | 9.3 | 187  | 2 | US-08-846-021A-5         | Sequence 5, Appli |
| 41 | 65.5 | 9.3 | 234  | 1 | US-08-366-783-4          | Sequence 4, Appli |
| 42 | 65.5 | 9.3 | 254  | 2 | US-08-767-026 <b>-</b> 7 | Sequence 7, Appli |
| 43 | 65.5 | 9.3 | 254  | 4 | US-09-319-275A-7         | Sequence 7, Appli |
| 44 | 65   | 9.2 | 964  | 4 | US-09-422-936-53         | Sequence 53, Appl |
| 45 | 64.5 | 9.1 | 614  | 1 | US-08-291-299-7          | Sequence 7, Appli |
|    |      |     |      |   |                          |                   |

## ALIGNMENTS

```
RESULT 1
US-08-700-607-1
; Sequence 1, Application US/08700607
; Patent No. 5858708
  GENERAL INFORMATION:
    APPLICANT: Bandman, Olga
    APPLICANT: Au-Young, Janice
    APPLICANT: Goli, Surya K.
    APPLICANT: Hillman, Jennifer L.
    TITLE OF INVENTION: TWO NOVEL HUMAN NSP-LIKE PROTEINS
    NUMBER OF SEQUENCES: 9
    CORRESPONDENCE ADDRESS:
      ADDRESSEE: Incyte Pharmaceuticals, Inc.
      STREET: 3174 Porter Drive
      CITY: Palo Alto
      STATE: CA
      COUNTRY: U.S.
      ZIP: 94304
```

```
MEDIUM TYPE: Diskette
;
      COMPUTER: IBM Compatible
      OPERATING SYSTEM: DOS
      SOFTWARE: FastSEQ Version 1.5
    CURRENT APPLICATION DATA:
      APPLICATION NUMBER: US/08/700,607
      FILING DATE: Filed Herewith
    ATTORNEY/AGENT INFORMATION:
      NAME: Billings, Lucy J.
      REGISTRATION NUMBER: 36,749
      REFERENCE/DOCKET NUMBER: PF-0114 US
    TELECOMMUNICATION INFORMATION:
      TELEPHONE: 415-855-0555
      TELEFAX: 415-845-4166
   INFORMATION FOR SEQ ID NO: 1:
    SEQUENCE CHARACTERISTICS:
      LENGTH: 199 amino acids
      TYPE: amino acid
      STRANDEDNESS: single
      TOPOLOGY: linear
    MOLECULE TYPE: peptide
    IMMEDIATE SOURCE:
      LIBRARY:
      CLONE: Consensus
US-08-700-607-1
  Query Match
                        71.3%; Score 503; DB 2; Length 199;
  Best Local Similarity 98.1%; Pred. No. 2.3e-52;
 Matches 102; Conservative 1; Mismatches 1; Indels
                                                            0; Gaps
                                                                       0;
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QУ
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             Db
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RESULT 2
US-08-700-607-7
; Sequence 7, Application US/08700607
; Patent No. 5858708
  GENERAL INFORMATION:
    APPLICANT: Bandman, Olga
    APPLICANT: Au-Young, Janice
    APPLICANT: Goli, Surya K.
    APPLICANT: Hillman, Jennifer L.
    TITLE OF INVENTION: TWO NOVEL HUMAN NSP-LIKE PROTEINS
    NUMBER OF SEQUENCES: 9
    CORRESPONDENCE ADDRESS:
      ADDRESSEE: Incyte Pharmaceuticals, Inc.
      STREET: 3174 Porter Drive
      CITY: Palo Alto
      STATE: CA
      COUNTRY: U.S.
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COMPUTER READABLE FORM:

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COMPUTER READABLE FORM:
      MEDIUM TYPE: Diskette
      COMPUTER: IBM Compatible
      OPERATING SYSTEM: DOS
      SOFTWARE: FastSEQ Version 1.5
     CURRENT APPLICATION DATA:
      APPLICATION NUMBER: US/08/700,607
      FILING DATE: Filed Herewith
    ATTORNEY/AGENT INFORMATION:
      NAME: Billings, Lucy J.
      REGISTRATION NUMBER: 36,749
      REFERENCE/DOCKET NUMBER: PF-0114 US
    TELECOMMUNICATION INFORMATION:
      TELEPHONE: 415-855-0555
      TELEFAX: 415-845-4166
   INFORMATION FOR SEQ ID NO: 7:
    SEQUENCE CHARACTERISTICS:
      LENGTH: 208 amino acids
      TYPE: amino acid
      STRANDEDNESS: single
      TOPOLOGY: linear
    MOLECULE TYPE: peptide
    IMMEDIATE SOURCE:
      LIBRARY: GenBank
      CLONE: 307311
US-08-700-607-7
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           9 KMDCVWSNWKSQAIDLLYWRDIKQTGIVFGSFLLLLFSLTQFSVVSVVAYLALAALSATI 68
          97 SFRIYKGVIQAIAKSDEGHPFRAYLESEVAISEELVQKYSN 137
QУ
             Db
          69 SFRIYKSVLQAVQKTDEGHPFKAYLELEITLSOEOIOKYTD 109
RESULT 3
US-08-700-607-8
; Sequence 8, Application US/08700607
; Patent No. 5858708
  GENERAL INFORMATION:
    APPLICANT: Bandman, Olqa
    APPLICANT: Au-Young, Janice
    APPLICANT: Goli, Surya K.
    APPLICANT: Hillman, Jennifer L.
    TITLE OF INVENTION: TWO NOVEL HUMAN NSP-LIKE PROTEINS
    NUMBER OF SEQUENCES: 9
    CORRESPONDENCE ADDRESS:
      ADDRESSEE: Incyte Pharmaceuticals, Inc.
      STREET: 3174 Porter Drive
      CITY: Palo Alto
      STATE: CA
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ZIP: 94304

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ZIP: 94304
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    CURRENT APPLICATION DATA:
      APPLICATION NUMBER: US/08/700,607
      FILING DATE: Filed Herewith
    ATTORNEY/AGENT INFORMATION:
      NAME: Billings, Lucy J.
      REGISTRATION NUMBER: 36,749
      REFERENCE/DOCKET NUMBER: PF-0114 US
    TELECOMMUNICATION INFORMATION:
      TELEPHONE: 415-855-0555
      TELEFAX: 415-845-4166
  INFORMATION FOR SEQ ID NO: 8:
    SEQUENCE CHARACTERISTICS:
      LENGTH: 267 amino acids
      TYPE: amino acid
      STRANDEDNESS: single
      TOPOLOGY: linear
    MOLECULE TYPE: peptide
    IMMEDIATE SOURCE:
      LIBRARY: GenBank
      CLONE: 281046
US-08-700-607-8
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Qу
             Db
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RESULT 4
US-08-700-607-6
; Sequence 6, Application US/08700607
; Patent No. 5858708
  GENERAL INFORMATION:
    APPLICANT: Bandman, Olga
    APPLICANT: Au-Young, Janice
    APPLICANT: Goli, Surya K.
    APPLICANT: Hillman, Jennifer L.
    TITLE OF INVENTION: TWO NOVEL HUMAN NSP-LIKE PROTEINS
    NUMBER OF SEQUENCES: 9
    CORRESPONDENCE ADDRESS:
      ADDRESSEE: Incyte Pharmaceuticals, Inc.
      STREET: 3174 Porter Drive
      CITY: Palo Alto
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COUNTRY: U.S.

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      APPLICATION NUMBER: US/08/700,607
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      FILING DATE: Filed Herewith
    ATTORNEY/AGENT INFORMATION:
      NAME: Billings, Lucy J.
      REGISTRATION NUMBER: 36,749
      REFERENCE/DOCKET NUMBER: PF-0114 US
    TELECOMMUNICATION INFORMATION:
      TELEPHONE: 415-855-0555
      TELEFAX: 415-845-4166
   INFORMATION FOR SEQ ID NO: 6:
    SEQUENCE CHARACTERISTICS:
      LENGTH: 356 amino acids
      TYPE: amino acid
      STRANDEDNESS: single
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    MOLECULE TYPE: peptide
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US-08-700-607-6
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Qу
         106 QAIAKSDEGHPFRAYLESEVAISEELVOKYSN 137
             Db
         226 QAVQKTDEGHPFKAYLELEITLSQEQIQKYTD 257
RESULT 5
US-08-700-607-5
; Sequence 5, Application US/08700607
; Patent No. 5858708
  GENERAL INFORMATION:
    APPLICANT: Bandman, Olga
    APPLICANT: Au-Young, Janice
    APPLICANT: Goli, Surya K.
    APPLICANT: Hillman, Jennifer L.
    TITLE OF INVENTION: TWO NOVEL HUMAN NSP-LIKE PROTEINS
    NUMBER OF SEQUENCES: 9
    CORRESPONDENCE ADDRESS:
      ADDRESSEE: Incyte Pharmaceuticals, Inc.
      STREET: 3174 Porter Drive
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STATE: CA

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STATE: CA
       COUNTRY: U.S.
       ZIP: 94304
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       COMPUTER: IBM Compatible
       OPERATING SYSTEM: DOS
       SOFTWARE: FastSEQ Version 1.5
     CURRENT APPLICATION DATA:
      APPLICATION NUMBER: US/08/700,607
      FILING DATE: Filed Herewith
    ATTORNEY/AGENT INFORMATION:
      NAME: Billings, Lucy J.
       REGISTRATION NUMBER: 36,749
       REFERENCE/DOCKET NUMBER: PF-0114 US
     TELECOMMUNICATION INFORMATION:
      TELEPHONE: 415-855-0555
      TELEFAX: 415-845-4166
   INFORMATION FOR SEQ ID NO: 5:
     SEQUENCE CHARACTERISTICS:
      LENGTH: 776 amino acids
      TYPE: amino acid
      STRANDEDNESS: single
      TOPOLOGY: linear
    MOLECULE TYPE: peptide
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      CLONE: 307307
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US-08-700-607-3
; Sequence 3, Application US/08700607
; Patent No. 5858708
  GENERAL INFORMATION:
    APPLICANT: Bandman, Olga
    APPLICANT: Au-Young, Janice
    APPLICANT: Goli, Surya K.
    APPLICANT: Hillman, Jennifer L.
    TITLE OF INVENTION: TWO NOVEL HUMAN NSP-LIKE PROTEINS
    NUMBER OF SEQUENCES: 9
    CORRESPONDENCE ADDRESS:
      ADDRESSEE: Incyte Pharmaceuticals, Inc.
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CITY: Palo Alto

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CITY: Palo Alto
      STATE: CA
      COUNTRY: U.S.
      ZIP: 94304
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      MEDIUM TYPE: Diskette
      COMPUTER: IBM Compatible
      OPERATING SYSTEM: DOS
      SOFTWARE: FastSEQ Version 1.5
    CURRENT APPLICATION DATA:
      APPLICATION NUMBER: US/08/700,607
      FILING DATE: Filed Herewith
    ATTORNEY/AGENT INFORMATION:
      NAME: Billings, Lucy J.
      REGISTRATION NUMBER: 36,749
      REFERENCE/DOCKET NUMBER: PF-0114 US
    TELECOMMUNICATION INFORMATION:
      TELEPHONE: 415-855-0555
      TELEFAX: 415-845-4166
   INFORMATION FOR SEQ ID NO:
    SEQUENCE CHARACTERISTICS:
      LENGTH: 241 amino acids
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      CLONE: 31870
US-08-700-607-3
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Qу
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RESULT 7
US-09-149-476-563
; Sequence 563, Application US/09149476
; Patent No. 6420526
; GENERAL INFORMATION:
  APPLICANT: Rosen et al.
  TITLE OF INVENTION: 186 Human Secreted proteins
  FILE REFERENCE: PZ002P1
  CURRENT APPLICATION NUMBER: US/09/149,476
  CURRENT FILING DATE: 1998-09-08
  EARLIER APPLICATION NUMBER: PCT/US98/04493
  EARLIER FILING DATE: 1998-03-06
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STREET: 3174 Porter Drive

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EARLIER FILING DATE: 1997-03-07
EARLIER APPLICATION NUMBER: 60/040,626
EARLIER FILING DATE: 1997-03-07
EARLIER APPLICATION NUMBER: 60/040,334
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EARLIER APPLICATION NUMBER: 60/040,336
EARLIER FILING DATE: 1997-03-07
EARLIER APPLICATION NUMBER: 60/040,163
EARLIER FILING DATE: 1997-03-07
EARLIER APPLICATION NUMBER: 60/047,600
EARLIER FILING DATE: 1997-05-23
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EARLIER APPLICATION NUMBER: 60/047,597
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EARLIER APPLICATION NUMBER: 60/047,581
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EARLIER APPLICATION NUMBER: 60/047,598
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EARLIER APPLICATION NUMBER: 60/047,582
EARLIER FILING DATE: 1997-05-23
EARLIER APPLICATION NUMBER: 60/047,596
EARLIER FILING DATE: 1997-05-23
EARLIER APPLICATION NUMBER: 60/047,612
EARLIER FILING DATE: 1997-05-23
EARLIER APPLICATION NUMBER: 60/047,632
EARLIER FILING DATE: 1997-05-23
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EARLIER APPLICATION NUMBER: 60/047,601

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Aizawa K., Izawa M., Nishi K., Kiyosawa H., Kondo S., Yamanaka I., Saito T., Okazaki Y., Gojobori T., Bono H., Kasukawa T., Saito R.,
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     Kadota K., Matsuda H.A., Ashburner M., Batalov S., Casavant T.,
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     Fleischmann W., Gaasterland T., Gissi C., King B., Kochiwa H.,
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     Schriml L.M., Staubli F., Suzuki R., Tomita M., Wagner L., Washio T.,
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     Sakai K., Okido T., Furuno M., Aono H., Baldarelli R., Barsh G.,
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     Blake J., Boffelli D., Bojunga N., Carninci P., de Bonaldo M.F.,
RA
     Brownstein M.J., Bult C., Fletcher C., Fujita M., Gariboldi M.,
RA
     Gustincich S., Hill D., Hofmann M., Hume D.A., Kamiya M., Lee N.H.,
     Lyons P., Marchionni L., Mashima J., Mazzarelli J., Mombaerts P.,
RA
     Nordone P., Ring B., Ringwald M., Rodriguez I., Sakamoto N.,
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     Sasaki H., Sato K., Schoenbach C., Seya T., Shibata Y., Storch K.-F.,
RA
RA
     Suzuki H., Toyo-oka K., Wang K.H., Weitz C., Whittaker C., Wilming L.,
     Wynshaw-Boris A., Yoshida K., Hasegawa Y., Kawaji H., Kohtsuki S.,
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     "Functional annotation of a full-length mouse cDNA collection.";
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         similarity).
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     -!- SUBCELLULAR LOCATION: Integral membrane protein. Anchored to the
CC
         membrane of the endoplasmic reticulum through 2 putative
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CC
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     -!- SIMILARITY: Contains 1 reticulon domain.
     CC
CC
     This SWISS-PROT entry is copyright. It is produced through a collaboration
     between the Swiss Institute of Bioinformatics and the EMBL outstation -
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CC
     the European Bioinformatics Institute. There are no restrictions on its
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     use by non-profit institutions as long as its content is in no way
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     modified and this statement is not removed. Usage by and for commercial
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     or send an email to license@isb-sib.ch).
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     GO; GO:0005635; C:nuclear membrane; ISS.
DR
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     GO; GO:0019987; P:negative regulation of anti-apoptosis; ISS.
DR
     GO; GO:0030517; P:negative regulation of axon extension; ISS.
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                                  CYTOPLASMIC (Potential).
FT
     TRANSMEM
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                                  POTENTIAL.
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Arakawa T., Hara A., Fukunishi Y., Konno H., Adachi J., Fukuda S.,

RA

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FT
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                       137
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                138
                       162
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                       199
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FT
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              Db
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RTN4 RAT
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AC
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DT
     28-FEB-2003 (Rel. 41, Created)
DT
     28-FEB-2003 (Rel. 41, Last sequence update)
DT
     15-SEP-2003 (Rel. 42, Last annotation update)
     Reticulon 4 (Neurite outgrowth inhibitor) (Nogo protein) (Foocen)
DE
     (Glut4 vesicle 20 kDa protein).
DE
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GN
OS
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OC
     Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi;
OC.
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OX
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RN
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     STRAIN=Sprague-Dawley; TISSUE=Adipocyte;
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RX
    Morris N.J., Ross S.A., Neveu J.M., Lane W.S., Lienhard G.E.;
RA
RТ
     "Cloning and characterization of a 22 kDa protein from rat adipocytes:
RT
     a new member of the reticulon family.";
RL
    Biochim. Biophys. Acta 1450:68-76(1999).
RN
RΡ
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RX
    MEDLINE=20129258; PubMed=10667796;
RA
     Chen M.S., Huber A.B., Van der Haar M.E., Frank M., Schnell L.,
RA
     Spillmann A.A., Christ F., Schwab M.E.;
RТ
     "Nogo-A is a myelin-associated neurite outgrowth inhibitor and an
RT
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RL
    Nature 403:434-439(2000).
RN
     [3]
RP
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     STRAIN-Wistar Kyoto; TISSUE-Vascular smooth muscle;
RC
RA
     Ito T., Schwartz S.M.;
RT
     "Cloning of a member of the reticulon gene family in rat: one of two
RT
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RN
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RX
     MEDLINE=22033691; PubMed=12037567;
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     GrandPre T., Li S., Strittmatter S.M.;
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     Nature 417:547-551(2002).
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CC
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     -!- SUBUNIT: Binds to RTN4R. Interacts with Bcl-xl and Bcl-2 (By
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         similarity).
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     -!- SUBCELLULAR LOCATION: Integral membrane protein. Anchored to the
         membrane of the endoplasmic reticulum through 2 putative
CC
CC
         transmembrane domains (By similarity).
CC
     -!- ALTERNATIVE PRODUCTS:
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         Event=Alternative splicing; Named isoforms=4;
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         Name=2; Synonyms=Nogo-B, Foocen-M1;
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           IsoId=Q9JK11-2; Sequence=VSP 005658;
CC
         Name=3; Synonyms=Nogo-C, VP20;
CC
           IsoId=Q9JK11-3; Sequence=VSP 005656, VSP_005657;
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         Name=4; Synonyms=Foocen-M2;
CC
           IsoId=Q9JK11-4; Sequence=VSP 005659;
CC
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CC
         nerve, spinal cord and cerebral cortex. Isoforms 1 and 2 are
CC
         present in dorsal root ganglion, sciatic nerve and PC12 cells
CC
         after longer exposure. Isoforms 2 and 3 are detected in kidney,
CC
         cartilage, skin, lung and spleen. Isoform 3 is expressed at high
         level in skeletal muscle. In adult animals isoform 1 is expressed
CC
CC
         mainly in the nervous system.
CC
     -!- SIMILARITY: Contains 1 reticulon domain.
CC
     CC
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     between the Swiss Institute of Bioinformatics and the EMBL outstation -
     the European Bioinformatics Institute. There are no restrictions on its
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     or send an email to license@isb-sib.ch).
CC
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    GO; GO:0005635; C:nuclear membrane; ISS.
DR
    GO; GO:0005515; F:protein binding activity; ISS.
    GO; GO:0019987; P:negative regulation of anti-apoptosis; ISS.
DR
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    GO; GO:0030517; P:negative regulation of axon extension; ISS.
    InterPro; IPR003388; Reticulon.
DR
DR
    Pfam; PF02453; Reticulon; 1.
DR
    PROSITE; PS50845; RETICULON; 1.
    Endoplasmic reticulum; Alternative splicing; Transmembrane.
KW
FT
    DOMAIN
                 1
                       989
                               CYTOPLASMIC (Potential).
FT
    TRANSMEM
                990
                      1010
                                 POTENTIAL.
```

RP

FUNCTION.

```
FT
                                 LUMENAL (Potential).
     DOMAIN
                1011
                       1104
FT
     TRANSMEM
                1105
                       1125
                                 POTENTIAL.
FT
     DOMAIN
                1126
                       1163
                                 CYTOPLASMIC (Potential).
FT
     DOMAIN
                 976
                       1163
                                 RETICULON.
FT
     DOMAIN
                  33
                        46
                                 POLY-GLU.
FТ
                  73
     DOMAIN
                        76
                                 POLY-ALA.
FT
     DOMAIN
                 140
                       145
                                 POLY-PRO.
FT
     VARSPLIC
                  1
                        964
                                 Missing (in isoform 3).
FT
                                 /FTId=VSP 005656.
FT
                                 AVLSAELSKTS -> MDGQKKHWKDK (in isoform
     VARSPLIC
                 965
                        975
FΤ
                                 3).
FT
                                 /FTId=VSP 005657.
FT
     VARSPLIC
                173
                                 Missing (in isoform 2).
                        975
FT
                                 /FTId=VSP 005658.
FT
     VARSPLIC
                192
                       975
                                 Missing (in isoform 4).
FT
                                 /FTId=VSP 005659.
FT
     CONFLICT
                1130
                       1131
                                 MISSING (IN REF. 3; AAD31020).
SO
     SEQUENCE
                1163 AA; 126386 MW; 8CB894B09E94F0B6 CRC64;
  Query Match
                         63.4%; Score 447; DB 1; Length 1163;
  Best Local Similarity
                         96.9%; Pred. No. 6.1e-36;
  Matches
                                0; Mismatches 3; Indels
          93; Conservative
                                                                0; Gaps
Qу
           46 KDKVVDLLYWRDIKKTGVVFGASLFLLLSLTVFSIVSVTAYIALALLSVTISFRIYKGVI 105
                 973 KTSVVDLLYWRDIKKTGVVFGASLFLLLSLTVFSIVSVTAYIALALLSVTISFRIYKGVI 1032
Db
Qу
          106 QAIAKSDEGHPFRAYLESEVAISEELVOKYSNSALG 141
              Db
         1033 QAIQKSDEGHPFRAYLESEVAISEELVQKYSNSALG 1068
RESULT 3
RTN4 HUMAN
ID
     RTN4 HUMAN
                   STANDARD;
                                  PRT; 1192 AA.
     Q9NQC3; O94962; Q9BXG5; Q9H212; Q9H3I3; Q9UQ42; Q9Y293; Q9Y2Y7;
AC
AC
     09Y5U6;
     28-FEB-2003 (Rel. 41, Created)
DT
DT
     28-FEB-2003 (Rel. 41, Last sequence update)
     15-SEP-2003 (Rel. 42, Last annotation update)
DT
ĎΕ
     Reticulon 4 (Neurite outgrowth inhibitor) (Nogo protein) (Foocen)
DE
     (Neuroendocrine-specific protein) (NSP) (Neuroendocrine specific
DE
     protein C homolog) (RTN-x) (Reticulon 5) (My043 protein).
     RTN4 OR NOGO OR ASY OR KIAA0886.
GN
OS
     Homo sapiens (Human).
OC.
     Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi;
     Mammalia; Eutheria; Primates; Catarrhini; Hominidae; Homo.
OC.
ΟX
     NCBI TaxID=9606;
RN
     [1]
RP
     SEQUENCE FROM N.A. (ISOFORMS 1; 2 AND 3).
RX
     MEDLINE=20129242; PubMed=10667780:
RA
     Prinjha R., Moore S.E., Vinson M., Blake S., Morrow R., Christie G.,
RA
     Michalovich D., Simmons D.L., Walsh F.S.;
RT
     "Inhibitor of neurite outgrowth in humans.";
RL
     Nature 403:383-384(2000).
RN
     SEQUENCE FROM N.A. (ISOFORMS 1 AND 2).
```

```
RC
     TISSUE=Brain;
     MEDLINE=21010696; PubMed=11126360;
RX
     Tagami S., Equchi Y., Kinoshita M., Takeda M., Tsujimoto Y.;
RA
     "A novel protein, RTN-XS, interacts with both Bcl-XL and Bcl-2 on
RT
RT
     endoplasmic reticulum and reduces their anti-apoptotic activity.";
RL
     Oncogene 19:5736-5746(2000).
RN
RP
     SEQUENCE FROM N.A. (ISOFORMS 1; 2 AND 3).
RX
     MEDLINE=20237542; PubMed=10773680;
RA
     Yang J., Yu L., Bi A.D., Zhao S.-Y.;
     "Assignment of the human reticulon 4 gene (RTN4) to chromosome
RT
RT
     2p14-->2p13 by radiation hybrid mapping.";
RL
     Cytogenet. Cell Genet. 88:101-102(2000).
RN
     SEQUENCE FROM N.A. (ISOFORM 4).
RΡ
RA
     Jin W.-L., Ju G.;
RT
     "Developmentally-regulated alternative splicing in a novel Nogo-A.";
     Submitted (NOV-2000) to the EMBL/GenBank/DDBJ databases.
RL
RN
RP
     SEQUENCE FROM N.A. (ISOFORMS 2 AND 3).
     TISSUE=Placenta, and Skeletal muscle;
RC
     Ito T., Schwartz S.M.;
RA
RT
     "Cloning of a member of the reticulon gene family in human.";
RL
     Submitted (FEB-1999) to the EMBL/GenBank/DDBJ databases.
RN
     SEQUENCE FROM N.A. (ISOFORM 2).
RP
     TISSUE=Fibroblast:
RC
     Yutsudo M.;
RA
RT
     "Isolation of a cell death-inducing gene.";
RL
     Submitted (JUN-1998) to the EMBL/GenBank/DDBJ databases.
RN
RΡ
     SEQUENCE FROM N.A. (ISOFORM 3).
RC
     TISSUE=Pituitary;
     Song H., Peng Y., Zhou J., Huang Q., Dai M., Mao Y.M., Yu Y., Xu X.,
RA
RA
     Luo B., Hu R., Chen J.;
RT
     "Human neuroendocrine-specific protein C (NSP) homolog gene.";
     Submitted (JUL-1998) to the EMBL/GenBank/DDBJ databases.
RL
RN
RP
     SEQUENCE FROM N.A. (ISOFORM 3).
     Gu J.R., Wan D.F., Zhao X.T., Zhou X.M., Jiang H.Q., Zhang P.P.,
RA
RA
     Qin W.X., Huang Y., Qiu X.K., Qian L.F., He L.P., Li H.N., Yu Y.,
RA
     Yu J., Han L.H.;
     "Novel human cDNA clone with function of inhibiting cancer cell
RT
RT
RL
     Submitted (AUG-1999) to the EMBL/GenBank/DDBJ databases.
RN
RΡ
     SEQUENCE FROM N.A. (ISOFORM 1).
RC
     TISSUE=Brain;
RX
     MEDLINE=99156230; PubMed=10048485;
RA
     Nagase T., Ishikawa K.-I., Suyama M., Kikuno R., Hirosawa M.,
     Miyajima N., Tanaka A., Kotani H., Nomura N., Ohara O.;
RA
RT
     "Prediction of the coding sequences of unidentified human genes. XII.
     The complete sequences of 100 new cDNA clones from brain which code
RT
RT
     for large proteins in vitro.";
     DNA Res. 5:355-364(1998).
RL
RN
     [10]
RΡ
     SEQUENCE FROM N.A. (ISOFORMS 2 AND 3).
```

```
TISSUE=Brain, Pancreas, Placenta, and Skeletal muscle;
RC
RX
     MEDLINE=22388257; PubMed=12477932;
     Strausberg R.L., Feingold E.A., Grouse L.H., Derge J.G.,
RA
     Klausner R.D., Collins F.S., Wagner L., Shenmen C.M., Schuler G.D.,
RA
RA
     Altschul S.F., Zeeberg B., Buetow K.H., Schaefer C.F., Bhat N.K.,
     Hopkins R.F., Jordan H., Moore T., Max S.I., Wang J., Hsieh F.,
RΑ
RA
     Diatchenko L., Marusina K., Farmer A.A., Rubin G.M., Hong L.,
RA
     Stapleton M., Soares M.B., Bonaldo M.F., Casavant T.L., Scheetz T.E.,
RA
     Brownstein M.J., Usdin T.B., Toshiyuki S., Carninci P., Prange C.,
     Raha S.S., Loquellano N.A., Peters G.J., Abramson R.D., Mullahy S.J.,
RA
RA
     Bosak S.A., McEwan P.J., McKernan K.J., Malek J.A., Gunaratne P.H.,
RΑ
     Richards S., Worley K.C., Hale S., Garcia A.M., Gay L.J., Hulyk S.W.,
RA
     Villalon D.K., Muzny D.M., Sodergren E.J., Lu X., Gibbs R.A.,
RA
     Fahey J., Helton E., Ketteman M., Madan A., Rodrigues S., Sanchez A.,
RA
     Whiting M., Madan A., Young A.C., Shevchenko Y., Bouffard G.G.,
RA
     Blakesley R.W., Touchman J.W., Green E.D., Dickson M.C.,
     Rodriguez A.C., Grimwood J., Schmutz J., Myers R.M.,
RΑ
     Butterfield Y.S.N., Krzywinski M.I., Skalska U., Smailus D.E.,
RA
RA
     Schnerch A., Schein J.E., Jones S.J.M., Marra M.A.;
RT
     "Generation and initial analysis of more than 15,000 full-length
RT
     human and mouse cDNA sequences.";
RL
     Proc. Natl. Acad. Sci. U.S.A. 99:16899-16903(2002).
RN
     [11]
RP
     SEQUENCE FROM N.A. (ISOFORM 3).
     MEDLINE=20499367; PubMed=11042152;
RX
RA
     Zhang Q.-H., Ye M., Wu X.-Y., Ren S.-X., Zhao M., Zhao C.-J., Fu G.,
RA
     Shen Y., Fan H.-Y., Lu G., Zhong M., Xu X.-R., Han Z.-G., Zhang J.-W.,
     Tao J., Huang Q.-H., Zhou J., Hu G.-X., Gu J., Chen S.-J., Chen Z.;
RТ
     "Cloning and functional analysis of cDNAs with open reading frames for
RT
     300 previously undefined genes expressed in CD34+ hematopoietic
RT
     stem/progenitor cells.";
RL
     Genome Res. 10:1546-1560(2000).
RN
     [12]
RΡ
     SEQUENCE OF 482-1192 FROM N.A. (ISOFORM 1/4).
RC
     TISSUE=Brain;
     Mao Y.M., Xie Y., Zheng Z.H.;
RA
RL
     Submitted (MAY-1998) to the EMBL/GenBank/DDBJ databases.
RN
RP
     SEQUENCE OF 186-1192 FROM N.A. (ISOFORM 1).
RC
     TISSUE=Testis;
RA
     Sha J.H., Zhou Z.M., Li J.M.;
RL
     Submitted (JAN-2001) to the EMBL/GenBank/DDBJ databases.
RN
     [14]
RΡ
     TOPOLOGY.
     TISSUE=Brain;
RC
RX
     MEDLINE=20129259; PubMed=10667797;
RA
     GrandPre T., Nakamura F., Vartanian T., Strittmatter S.M.;
RT
     "Identification of the Nogo inhibitor of axon regeneration as a
RT
     Reticulon protein.";
RL
     Nature 403:439-444(2000).
RN
     [15]
RΡ
     FUNCTION.
RC
     TISSUE=Brain;
RX
     MEDLINE=21069055; PubMed=11201742;
RA
     Fournier A.E., Grandpre T., Strittmatter S.M.;
RT
     "Identification of a receptor mediating Nogo-66 inhibition of axonal
RT
     regeneration.";
```

```
Nature 409:341-346(2001).
RL
RN
     [16]
RP
    REVIEW.
RX
    MEDLINE=21888956; PubMed=11891768;
RA
    Ng C.E.L., Tang B.L.;
     "Nogos and the Nogo-66 receptor: factors inhibiting CNS neuron
RT
    regeneration.";
RT
    J. Neurosci. Res. 67:559-565(2002).
RL
CC
     -!- FUNCTION: Potent neurite outgrowth inhibitor which may also help
        block the regeneration of the nervous central system in adults.
CC
        Isoform 2 reduces the anti-apoptotic activity of Bcl-xl and Bcl-2.
CC
        This is likely consecutive to their change in subcellular
CC
CC
        location, from the mitochondria to the endoplasmic reticulum,
CC
        after binding and sequestration.
CC
     -!- SUBUNIT: Binds to RTN4R. Interacts with Bcl-xl and Bcl-2.
CC
     -!- SUBCELLULAR LOCATION: Integral membrane protein. Endoplasmic
CC
        reticulum. Anchored to the membrane of the endoplasmic reticulum
CC
        through 2 putative transmembrane domains.
CC
     -!- ALTERNATIVE PRODUCTS:
CC
        Event=Alternative splicing; Named isoforms=4;
CC
        Name=1; Synonyms=RTN 4A, Nogo-A, RTN-xL;
CC
          IsoId=Q9NQC3-1; Sequence=Displayed;
CC
        Name=2; Synonyms=RTN 4B, Nogo-B, RTN-xS, Foocen-M;
CC
          IsoId=Q9NQC3-2; Sequence=VSP 005655;
CC
        Name=3; Synonyms=RTN 4C, Nogo-C, Foocen-S;
          IsoId=Q9NQC3-3; Sequence=VSP 005652, VSP 005653;
CC
CC
CC
          IsoId=Q9NQC3-4; Sequence=VSP 005654;
CC
     -!- TISSUE SPECIFICITY: Isoform 1 is specifically expressed in brain
CC
        and testis and weakly in heart and skeletal muscle. Isoform 2 is
CC
        widely expressed excepted for the liver. Isoform 3 is expressed in
CC
        brain, skeletal muscle and adipocytes. Isoform 4 is testis-
CC
        specific.
CC
     -!- SIMILARITY: Contains 1 reticulon domain.
CC
     -!- CAUTION: Ref.11 sequence differs from that shown due to
CC
        frameshifts in positions 1149 and 1156.
CC
     _____
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     between the Swiss Institute of Bioinformatics and the EMBL outstation -
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     the European Bioinformatics Institute. There are no restrictions on its
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     use by non-profit institutions as long as its content is in no way
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     modified and this statement is not removed. Usage by and for commercial
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     entities requires a license agreement (See http://www.isb-sib.ch/announce/
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     or send an email to license@isb-sib.ch).
CC
     EMBL; AJ251383; CAB99248.1; -.
DR
DR
     EMBL; AJ251384; CAB99249.1; -.
DR
     EMBL; AJ251385; CAB99250.1; -.
DR
     EMBL; AB040462; BAB18927.1; -.
DR
     EMBL; AB040463; BAB18928.1; -.
     EMBL; AF148537; AAG12176.1; -.
DR
DR
     EMBL; AF148538; AAG12177.1; -.
DR
     EMBL; AF087901; AAG12205.1; -.
DR
     EMBL; AF320999; AAG40878.1; -.
DR
     EMBL; AF132047; AAD31021.1; -.
DR
     EMBL; AF132048; AAD31022.1; -.
     EMBL; AB015639; BAA83712.1; -.
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EMBL; AF077050; AAD27783.1; -.
DR
DR
    EMBL; AF177332; AAG17976.1; -.
DR
    EMBL; AB020693; BAA74909.1; -.
DR
    EMBL; BC001035; AAH01035.1; -.
DR
    EMBL; BC007109; AAH07109.1; -.
    EMBL; BC014366; AAH14366.1; -.
DR
                         63.4%; Score 447; DB 1; Length 1192;
 Query Match
 Best Local Similarity
                         96.9%; Pred. No. 6.2e-36;
           93; Conservative
                             0; Mismatches
                                                 3; Indels
                                                               0; Gaps
                                                                           0;
Qу
          46 KDKVVDLLYWRDIKKTGVVFGASLFLLLSLTVFSIVSVTAYIALALLSVTISFRIYKGVI 105
                1002 KTSVVDLLYWRDIKKTGVVFGASLFLLLSLTVFSIVSVTAYIALALLSVTISFRIYKGVI 1061
Dh
Qу
         106 QAIAKSDEGHPFRAYLESEVAISEELVQKYSNSALG 141
             Db
        1062 QAIQKSDEGHPFRAYLESEVAISEELVQKYSNSALG 1097
RESULT 4
RTN1 HUMAN
    RTN1 HUMAN
                   STANDARD;
                                  PRT;
                                        776 AA.
AC
    016799; 016800; 016801;
DT
    16-OCT-2001 (Rel. 40, Created)
DT
    16-OCT-2001 (Rel. 40, Last sequence update)
DT
    15-SEP-2003 (Rel. 42, Last annotation update)
    Reticulon 1 (Neuroendocrine-specific protein).
GN
    RTN1 OR NSP.
OS
    Homo sapiens (Human).
OC
     Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi;
OC
    Mammalia; Eutheria; Primates; Catarrhini; Hominidae; Homo.
OX
    NCBI TaxID=9606;
RN
    [1]
RΡ
     SEQUENCE FROM N.A. (ISOFORMS RTN1-A; RTN1-B AND RTN1-C).
RC
    TISSUE=Lung carcinoma;
    MEDLINE=93293865; PubMed=7685762;
RX
RA
    Roebroek A.J.M., Van de Velde H.J.K., Van Bokhoven A., Broers J.L.V.,
RA
    Ramaekers F.C.S., Van de Ven W.J.M.;
RT
     "Cloning and expression of alternative transcripts of a novel
RT
     neuroendocrine-specific gene and identification of its 135-kDa
RT
     translational product.";
RL
    J. Biol. Chem. 268:13439-13447(1993).
RN
     [2]
RP
    ALTERNATIVE SPLICING.
RX
    MEDLINE=96429995; PubMed=8833145;
RA
    Roebroek A.J.M., Ayoubi T.A.Y., Van de Velde H.J.K.,
RA
    Schoenmakers E.F.P.M., Pauli I.G.L., Van de Ven W.J.M.;
RT
     "Genomic organization of the human NSP gene, prototype of a novel gene
RT
     family encoding reticulons.";
    Genomics 32:191-199(1996).
RL
RN
     [3]
RΡ
    TISSUE SPECIFICITY.
RX
    MEDLINE=98228245; PubMed=9560466;
RA
    Hens J., Nuydens R., Geerts H., Senden N.H., Van de Ven W.J.M.,
    Roebroek A.J., van de Velde H.J., Ramaekers F.C., Broers J.L.;
RA
RT
     "Neuronal differentiation is accompanied by NSP-C expression.";
```

```
RL
    Cell Tissue Res. 292:229-237(1998).
    -!- FUNCTION: MAY BE INVOLVED IN NEUROENDOCRINE SECRETION OR IN
CC
CC
        MEMBRANE TRAFFICKING IN NEUROENDOCRINE CELLS.
CC
     -!- SUBCELLULAR LOCATION: Endoplasmic reticulum membrane.
CC
    -!- ALTERNATIVE PRODUCTS:
CC
        Event=Alternative splicing; Named isoforms=3;
CC
        Name=RTN1-A; Synonyms=NSP-A;
CC
          IsoId=Q16799-1; Sequence=Displayed;
CC
        Name=RTN1-B; Synonyms=NSP-B;
          IsoId=Q16799-2; Sequence=VSP 005644;
CC
CC
        Name=RTN1-C; Synonyms=NSP-C;
CC
          IsoId=Q16799-3; Sequence=VSP 005645, VSP 005646;
CC
    -!- TISSUE SPECIFICITY: EXPRESSED IN NEURAL AND NEUROENDOCRINE TISSUES
CC
        AND CELL CULTURES DERIVED THEREFROM. EXPRESSION OF ISOFORM RTN1-C
        IS STRONGLY CORRELATED WITH NEURONAL DIFFERENTIATION.
CC
CC
     -!- PTM: ISOFORMS RTN1-A AND RTN5以中26以由
                                                HORYLATED.
CC
     -!- SIMILARITY: Contains 1 reticulon domain.
CC
     ______
CC
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CC
DR
    EMBL; L10333; AAA59950.1; -.
DR
    EMBL; L10334; AAA59951.1; -.
    EMBL; L10335; AAA59952.1; -.
DR
DR
    PIR; A46583; A46583.
           0904; I609p4475X
DR
    PIR;
    Genew; HGNC: 10467; RTN1.
DR
DR
    MIM; 600865; -.
DR
    GO; GO:0030176; C:endoplasmic reticulum membrane, intrinsic p. . .; TAS.
DR
    GO; GO:0004871; F:signal transducer activity; NAS.
DR
    GO; GO:0030182; P:neuron differentiation; TAS.
DR
    GO; GO:0007165; P:signal transduction; NAS.
DR
    InterPro; IPR003388; Reticulon.
DR
    Pfam; PF02453; Reticulon; 1.
DR
    PROSITE; PS50845; RETICULON; 1.
KW
    Endoplasmic reticulum; Alternative splicing; Transmembrane;
KW
    Phosphorylation.
FT
    TRANSMEM
               603
                       623
                                POTENTIAL.
FT
    TRANSMEM
                726
                      746
                                POTENTIAL.
FT
    DOMAIN
                589 776
                                RETICULON.
                609
FT
    DOMAIN
                      612
                                POLY-LEU.
FT
    VARSPLIC
               1
                      420
                                Missing (in isoform RTN1-B).
FT
                                /FTId=VSP 005644.
FT
    VARSPLIC
                1 568
                                Missing (in isoform RTN1-C).
FT
                                /FTId=VSP 005645.
FT
    VARSPLIC
                569
                       588
                                GPGPLGPGAPPPLLFLNKOK -> MOATADSTKMDCVWSNW
FT
                                KSQ (in isoform RTN1-C).
FT
                                /FTId=VSP 005646.
    SEQUENCE 776 AA; 83617 MW; CA5B6232353096FE CRC64;
SQ
 Query Match
                        47.8%; Score 337; DB 1; Length 776;
 Best Local Similarity 67.4%; Pred. No. 2.4e-25;
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Matches 62; Conservative 16; Mismatches 14; Indels
                                                             0; Gaps
                                                                          0;
          46 KDKVVDLLYWRDIKKTGVVFGASLFLLLSLTVFSIVSVTAYIALALLSVTISFRIYKGVI 105
Qу
             Db
         586 KQKAIDLLYWRDIKQTGIVFGSFLLLLFSLTQFSVVSVVAYLALAALSATISFRIYKSVL 645
         106 QAIAKSDEGHPFRAYLESEVAISEELVQKYSN 137
Qу
             646 QAVQKTDEGHPFKAYLELEITLSQEQIQKYTD 677
RESULT 5
RTN1 RAT
ΙD
    RTN1 RAT
                   STANDARD; PRT; 777 AA.
AC
    Q64548; Q64547;
    16-OCT-2001 (Rel. 40, Created)
    16-OCT-2001 (Rel. 40, Last sequence update)
DΤ
    15-SEP-2003 (Rel. 42, Last annotation update)
DT
DΕ
    Reticulon 1 (Neuroendocrine-specific protein) (S-rex).
GN
    RTN1 OR NSP.
OS
    Rattus norvegicus (Rat).
OC
    Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi;
OC
    Mammalia; Eutheria; Rodentia; Sciurognathi; Muridae; Murinae; Rattus.
OX
    NCBI TaxID=10116;
RN
     [1]
RP
    SEQUENCE FROM N.A. (ISOFORMS RTN1-B AND RTN1-S).
    STRAIN=Wistar; TISSUE=Brain cortex;
RC
RX
    MEDLINE=96386034; PubMed=8793864;
RA
    Baka I.D., Ninkina N.N., Pinon L.G.P., Adu J., Davies A.M.,
RA
    Georgiev G.P., Buchman V.L.;
RT
     "Intracellular compartmentalization of two differentially spliced s-
RT
    rex/NSP mRNAs in neurons.";
RL
    Mol. Cell. Neurosci. 7:289-303(1996).
    -!- FUNCTION: MAY BE INVOLVED IN NEUROENDOCRINE SECRETION OR IN
CC
CC
        MEMBRANE TRAFFICKING IN NEUROENDOCRINE CELLS.
CC
    -!- SUBCELLULAR LOCATION: ENDOPLASMIC RETICULUM MEMBRANE (BY
CC
        SIMILARITY).
    -!- ALTERNATIVE PRODUCTS:
CC
CC
        Event=Alternative splicing; Named isoforms=2;
CC
        Name=RTN1-B; Synonyms=S-RexB;
CC
          IsoId=Q64548-1; Sequence=Displayed;
CC
        Name=RTN1-S; Synonyms=S-RexS;
CC
          IsoId=Q64548-2; Sequence=VSP 005647, VSP 005648;
CC
    -!- TISSUE SPECIFICITY: EXPRESSED PREDOMINANTLY IN CENTRAL AND
CC
        PERIPHERAL NERVOUS SYSTEM OF NEWBORN AND ADULT RATS. LOW LEVELS
CC
        HAVE BEEN ALSO DETECTED IN HEART, ADRENAL GLAND AND SPLEEN.
CC
        EXPRESSION OF ISOFORM RTN1-B IS RESTRICTED TO PARTICULAR NEURONAL
CC
        TYPES.
CC
    -!- DEVELOPMENTAL STAGE: DETECTED ON EMBRYONIC DAY E10 IN THE
CC
        HINDBRAIN AND IN Ell IN THE FOREBRAIN. DURING THE NEXT 3 EMBRYONIC
CC
        DAYS THE LEVELS OF S-REXS INCREASES AND REMAINS STABLE AT E13 IN
        THE HINDBRAIN AND AT E14 IN THE FOREBRAIN. THE LEVELS OF S-REXB
CC
CC
        DOES NOT CHANGE AS SIGNIFICANTLY DURING DEVELOPMENT OF THE
CC
        HINDBRAIN.
CC
    -!- SIMILARITY: Contains 1 reticulon domain.
CC
```

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    or send an email to license@isb-sib.ch).
CC
CC
    _____
    EMBL; U17604; AAC53046.1; -.
DR
    EMBL; U17603; AAC53045.1; -.
DR
    InterPro; IPR003388; Reticulon.
DR
DR
    Pfam; PF02453; Reticulon; 1.
DR
    PROSITE; PS50845; RETICULON; 1.
    Endoplasmic reticulum; Alternative splicing; Transmembrane.
KW
FT
    TRANSMEM
             604 624 POTENTIAL.
    TRANSMEM
              727 747
                             POTENTIAL.
FΤ
FΤ
    DOMAIN
              590 777
                           RETICULON.
    DOMAIN
    DOMAIN 610 613
VARSPLIC 1 569
FT
                             POLY-LEU.
FT
                              Missing (in isoform RTN1-S).
FT
                              /FTId=VSP 005647.
FT
    VARSPLIC 570 589
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                              KSQ (in isoform RTN1-S).
FT
FT
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    SEQUENCE 777 AA; 83001 MW; AF7479C50F28D0AC CRC64;
SO
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 Best Local Similarity 67.4%; Pred. No. 2.4e-25;
 Matches 62; Conservative 16; Mismatches 14; Indels
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                                                                     0;
         46 KDKVVDLLYWRDIKKTGVVFGASLFLLLSLTVFSIVSVTAYIALALLSVTISFRIYKGVI 105
Qу
            587 KQKAIDLLYWRDIKQTGIVFGSFLLLLFSLTQFSVVSVVAYLALAALSATISFRIYKSVL 646
Db
         106 QAIAKSDEGHPFRAYLESEVAISEELVOKYSN 137
Ov
            647 QAVQKTDEGHPFKAYLELEITLSQEQIQKYTD 678
Dh
RESULT 6
RTN3 HUMAN
    RTN3 HUMAN
                 STANDARD; PRT; 236 AA.
AC
    095197;
DT
    16-OCT-2001 (Rel. 40, Created)
    16-OCT-2001 (Rel. 40, Last sequence update)
DΤ
    15-SEP-2003 (Rel. 42, Last annotation update)
    Reticulon protein 3 (Neuroendocrine-specific protein-like 2) (NSP-like
DE
DE
    protein II) (NSPLII).
    RTN3 OR NSPL2.
GN
OS
    Homo sapiens (Human).
OC
    Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi;
    Mammalia; Eutheria; Primates; Catarrhini; Hominidae; Homo.
OC
OX
    NCBI TaxID=9606;
RN
RP
    SEQUENCE FROM N.A., AND TISSUE SPECIFICITY.
RC
    TISSUE=Retina;
RX
    MEDLINE=99265974; PubMed=10331947;
    Moreira E.F., Jaworski C.J., Rodriguez I.R.;
RA
    "Cloning of a novel member of the reticulon gene family (RTN3): gene
RТ
```

between the Swiss Institute of Bioinformatics and the EMBL outstation -

```
CC
        reticulum (Potential).
CC
     -!- TISSUE SPECIFICITY: WIDELY EXPRESSED WITH HIGHEST EXPRESSION IN
CC
        BRAIN. THREE TIMES MORE ABUNDANT IN MACULA THAN IN PERIPHERAL
CC
        RETINA.
CC
    -!- SIMILARITY: Contains 1 reticulon domain.
CC
     ------
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CC
DR
    EMBL; AF059524; AAC99319.1; -.
DR
    EMBL; AF059529; AAD20951.1; -.
DR
    EMBL; AF059525; AAD20951.1; JOINED.
DR
    EMBL; AF059526; AAD20951.1; JOINED.
DR
    EMBL; AF059527; AAD20951.1; JOINED.
DR
    EMBL; AF059528; AAD20951.1; JOINED.
DR
    EMBL; AF119297; AAD26810.1; -.
DR
    EMBL; BC000634; AAH00634.1; -.
    EMBL; BC010556; AAH10556.1; -.
DR
DR
    EMBL; BC011394; AAH11394.1; -.
```

structure and chromosomal localization to 11g13.";

"Cloning and expression analysis of a cDNA encoding a novel

neuroendocrine-specific protein-like protein 1: NSPL1."; Submitted (JAN-1999) to the EMBL/GenBank/DDBJ databases.

Strausberg R.L., Feingold E.A., Grouse L.H., Derge J.G.,

Klausner R.D., Collins F.S., Wagner L., Shenmen C.M., Schuler G.D.,

Stapleton M., Soares M.B., Bonaldo M.F., Casavant T.L., Scheetz T.E.,

Raha S.S., Loquellano N.A., Peters G.J., Abramson R.D., Mullahy S.J.,

Richards S., Worley K.C., Hale S., Garcia A.M., Gay L.J., Hulyk S.W.,

Fahey J., Helton E., Ketteman M., Madan A., Rodrigues S., Sanchez A.,

Bosak S.A., McEwan P.J., McKernan K.J., Malek J.A., Gunaratne P.H.,

Altschul S.F., Zeeberg B., Buetow K.H., Schaefer C.F., Bhat N.K.,

Brownstein M.J., Usdin T.B., Toshiyuki S., Carninci P., Prange C.,

Hopkins R.F., Jordan H., Moore T., Max S.I., Wang J., Hsieh F.,

Diatchenko L., Marusina K., Farmer A.A., Rubin G.M., Hong L.,

Villalon D.K., Muzny D.M., Sodergren E.J., Lu X., Gibbs R.A.,

Blakesley R.W., Touchman J.W., Green E.D., Dickson M.C.,

Rodriguez A.C., Grimwood J., Schmutz J., Myers R.M.,

Schnerch A., Schein J.E., Jones S.J.M., Marra M.A.;

Proc. Natl. Acad. Sci. U.S.A. 99:16899-16903(2002).

human and mouse cDNA sequences.";

Whiting M., Madan A., Young A.C., Shevchenko Y., Bouffard G.G.,

Butterfield Y.S.N., Krzywinski M.I., Skalska U., Smailus D.E.,

"Generation and initial analysis of more than 15,000 full-length

-!- SUBCELLULAR LOCATION: Integral membrane protein. Endoplasmic

Huang X., Zhou Y., Du G., Yuan J., Qiang B.;

Genomics 58:73-81(1999).

SEQUENCE FROM N.A.

SEQUENCE FROM N.A.

TISSUE=Brain, Eye, and Lymph; MEDLINE=22388257; PubMed=12477932;

RT

RL

RN RP

RA

RT

RT

RL RN RP

RC

RX RA

RA RA

RA RA

RA

RA

RA

RT

RT

RL

```
EMBL; BC022993; AAH22993.1; -.
DR
DR
    Genew; HGNC:10469; RTN3.
DR
    MIM; 604249; -.
    GO; GO:0005615; C:extracellular space; TAS.
DR
    InterPro; IPR003388; Reticulon.
DR
    Pfam; PF02453; Reticulon; 1.
DR
    PROSITE; PS50845; RETICULON; 1.
DR
KW
    Transmembrane; Endoplasmic reticulum.
                               POTENTIAL.
FT
    TRANSMEM
                68
                      88
FT
    TRANSMEM
               177
                      197
                               POTENTIAL.
FT
    DOMAIN
                48
                      236
                               RETICULON.
    SEQUENCE 236 AA; 25609 MW; DDC6A4544ABCDFB7 CRC64;
SO
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 Best Local Similarity 60.9%; Pred. No. 3.7e-23;
          56; Conservative 18; Mismatches 18; Indels
 Matches
                                                             0; Gaps
                                                                        0;
          49 VVDLLYWRDIKKTGVVFGASLFLLLSLTVFSIVSVTAYIALALLSVTISFRIYKGVIQAI 108
Qу
             48 VHDLIFWRDVKKTGFVFGTTLIMLLSLAAFSVISVVSYLILALLSVTISFRIYKSVIQAV 107
Db
         109 AKSDEGHPFRAYLESEVAISEELVQKYSNSAL 140
QУ
              ||:||||:||:||:
         108 QKSEEGHPFKAYLDVDITLSSEAFHNYMNAAM 139
Db
RESULT 7
RTN3 MOUSE
    RTN3 MOUSE
                  STANDARD:
                                PRT:
                                       237 AA.
AC
    Q9ES97;
    16-OCT-2001 (Rel. 40, Created)
DT
    16-OCT-2001 (Rel. 40, Last sequence update)
DT
    28-FEB-2003 (Rel. 41, Last annotation update)
DT
DE
    Reticulon protein 3.
GN
    RTN3.
OS
    Mus musculus (Mouse).
OC
    Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi;
OC
    Mammalia; Eutheria; Rodentia; Sciurognathi; Muridae; Murinae; Mus.
OX
    NCBI TaxID=10090;
RN
    [1]
RP
    SEQUENCE FROM N.A.
RA
    Huang X., Zhou Y., Qiang H., Yuan J., Qiang B.;
RT
    "Cloning and expression profile of a novel mouse cDNA encoding a human
RT
    RTN3 homolog.";
RL
    Submitted (OCT-1999) to the EMBL/GenBank/DDBJ databases.
CC
    -! SUBCELLULAR LOCATION: Integral membrane protein. Endoplasmic
CC
        reticulum (Potential).
    -!- SIMILARITY: Contains 1 reticulon domain.
CC
CC
    ______
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DR
     MGD; MGI:1339970; Rtn3.
     InterPro; IPR003388; Reticulon.
DR
DR
     Pfam; PF02453; Reticulon; 1.
DR
     PROSITE; PS50845; RETICULON; 1.
KW
     Transmembrane; Endoplasmic reticulum.
FΤ
     TRANSMEM
                 69
                        89
                                 POTENTIAL.
FT
     TRANSMEM
                167
                       187
                                 POTENTIAL.
FT
                       237
     DOMATN
                 49
                                 RETICULON.
               237 AA; 25428 MW; EB60A0A7AC45F0DE CRC64;
SQ
     SEQUENCE
  Query Match
                         43.7%; Score 308; DB 1; Length 237;
  Best Local Similarity
                         59.8%; Pred. No. 4.6e-23;
  Matches
           55; Conservative
                              19; Mismatches
                                                18; Indels
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           49 VVDLLYWRDIKKTGVVFGASLFLLLSLTVFSIVSVTAYIALALLSVTISFRIYKGVIQAI 108
Qу
              | ||::||:||||||
Db
           49 VHDLIFWRDVKKTGFVFGTTLIMLLSLAAFSVISVVSYLILALLSVTISFRVYKSVIOAV 108
Qу
          109 AKSDEGHPFRAYLESEVAISEELVOKYSNSAL 140
               Db
          109 QKSEEGHPFKAYLDVDITLSSEAFHNYMNAAM 140
RESULT 8
RTN2 HUMAN
                                  PRT;
ID
     RTN2 HUMAN
                   STANDARD;
                                         545 AA.
AC
     075298; 060509;
DT
     16-OCT-2001 (Rel. 40, Created)
     16-OCT-2001 (Rel. 40, Last sequence update)
DT
     15-SEP-2003 (Rel. 42, Last annotation update)
DE
     Reticulon protein 2 (Neuroendocrine-specific protein-like 1) (NSP-like
DE
     protein 1) (NSPLI).
GN
     RTN2 OR NSPL1.
OS
     Homo sapiens (Human).
OC
     Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi;
OC
     Mammalia; Eutheria; Primates; Catarrhini; Hominidae; Homo.
OX
     NCBI TaxID=9606;
RN
     [1]
RP
     SEQUENCE FROM N.A., ALTERNATIVE SPLICING, AND TISSUE SPECIFICITY.
RC
     TISSUE=Lung carcinoma;
RX
     MEDLINE=98360096; PubMed=9693037;
RA
     Roebroek A.J.M., Contreras B., Pauli I.G.L., Van de Ven W.J.M.;
RT
     "cDNA cloning, genomic organization, and expression of the human RTN2
RT
     gene, a member of a gene family encoding reticulons.";
RL
     Genomics 51:98-106(1998).
RN
     [2]
RΡ
     SEQUENCE OF 108-545 FROM N.A. (ISOFORM RTN2-B).
RC
     TISSUE=Brain;
RX
    MEDLINE=98191726; PubMed=9530622;
RA
    Geisler J.G., Stubbs L.J., Wasserman W.W., Mucenski M.L.;
RT
     "Molecular cloning of a novel mouse gene with predominant muscle and
RT
    neural expression.";
RL
    Mamm. Genome 9:274-282(1998).
CC
    -!- SUBCELLULAR LOCATION: Integral membrane protein. Endoplasmic
CC
        reticulum (Potential).
CC
     -!- ALTERNATIVE PRODUCTS:
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DR

EMBL; AF195940; AAG31360.1; -.

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CC
        Event=Alternative splicing; Named isoforms=2;
CC
        Name=RTN2-A;
          IsoId=075298-1; Sequence=Displayed;
CC
          Note=Isoform RTN2-C is produced by alternative initiation at
CC
CC
          Met-341 of isoform RTN2-A;
CC
        Name=RTN2-B;
CC
          IsoId=075298-2; Sequence=VSP 005649;
CC
        Event=Alternative initiation;
CC
          Comment=2 isoforms, RTN2-A (shown here) and RTN2-C, are produced
          by alternative initiation at Met-1 and Met-341;
CC
CC
    -!- TISSUE SPECIFICITY: ISOFORM RTN2-C IS HIGHLY EXPRESSED IN SKELETAL
        MUSCLE.
CC
    -!- SIMILARITY: Contains 1 reticulon domain.
CC
CC
    ______
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CC
    ______
DR
    EMBL; AF004222; AAC32542.1; -.
    EMBL; AF004223; AAC32543.1; -.
DR
DR
    EMBL; AF004224; AAC32544.1; -.
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    EMBL; AF038540; AAC14910.1; -.
DR
    Genew; HGNC:10468; RTN2.
DR
    MIM; 603183; -.
    GO; GO:0030176; C:endoplasmic reticulum membrane, intrinsic p. . .; NAS.
    GO; GO:0004871; F:signal transducer activity; NAS.
DR
    GO; GO:0007165; P:signal transduction; NAS.
DR
DR
    InterPro; IPR003388; Reticulon.
    Pfam; PF02453; Reticulon; 1.
DR
    PROSITE; PS50845; RETICULON; 1.
DR
KW
    Endoplasmic reticulum; Alternative splicing; Transmembrane;
KW
    Alternative initiation.
FT
    CHAIN
                1
                      545
                               RETICULON PROTEIN 2, ISOFORM RTN2-A.
FT
    CHAIN
               341
                      545
                               RETICULON PROTEIN 2, ISOFORM RTN2-C.
    INIT MET
FT
               341
                      341
                               FOR ISOFORM RTN2-C.
FT
    TRANSMEM
               368
                   388
                               POTENTIAL.
FT
    TRANSMEM
               463
                     483
                               POTENTIAL.
FT
               345 545
    DOMAIN
                               RETICULON.
FT
    VARSPLIC 272 344
                               Missing (in isoform RTN2-B).
FT
                               /FTId=VSP 005649.
SO
    SEQUENCE 545 AA; 59263 MW; 971FD2F909E1E9E6 CRC64;
 Ouery Match
                       30.4%; Score 214; DB 1; Length 545;
 Best Local Similarity 46.7%; Pred. No. 1.8e-13;
         42; Conservative 21; Mismatches 27; Indels
                                                         0; Gaps
                                                                       0;
Qу
          48 KVVDLLYWRDIKKTGVVFGASLFLLLSLTVFSIVSVTAYIALALLSVTISFRIYKGVIQA 107
             344 KVADLLYWKDTRTSGVVFTGLMVSLLCLLHFSIVSVAAHLALLLLCGTISLRVYRKVLQA 403
Db
QУ
         108 IAKSDEGHPFRAYLESEVAISEELVOKYSN 137
            :: | :||:|||: :: :: | :: |:
Dh
         404 VHRGDGANPFQAYLDVDLTLTREQTERLSH 433
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RESULT 9
RTN2_MOUSE
    RTN2 MOUSE
                 STANDARD;
                                 PRT;
ID
                                        471 AA.
AC
     070622; 070620;
DT
     16-OCT-2001 (Rel. 40, Created)
DT
     16-OCT-2001 (Rel. 40, Last sequence update)
DT
     28-FEB-2003 (Rel. 41, Last annotation update)
     Reticulon protein 2 (Neuroendocrine-specific protein-like 1) (NSP-like
DE
_{
m DE}
     protein 1) (NSPLI).
GN
    RTN2 OR NSPL1.
OS
    Mus musculus (Mouse).
     Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi;
OC
OC
    Mammalia; Eutheria; Rodentia; Sciuroqnathi; Muridae; Murinae; Mus.
OX
    NCBI TaxID=10090;
RN
     [1]
RP
     SEQUENCE FROM N.A. (ISOFORMS 1 AND 2), AND TISSUE SPECIFICITY.
RC
     STRAIN=FVB/N, and 129/Sv; TISSUE=Cerebellum, and Skeletal muscle;
RX
    MEDLINE=98191726; PubMed=9530622;
RA
    Geisler J.G., Stubbs L.J., Wasserman W.W., Mucenski M.L.;
RT
     "Molecular cloning of a novel mouse gene with paredomsnent b
RT
    neural expression.";
    Mamm. Genome 9:274-282(1998).
RL
CC
     -!- SUBCELLULAR LOCATION: Membrane-bound. Endoplasmic reticulum
CC
        (Potential).
    -!- ALTERNATIVE PRODUCTS:
CC
        Event=Alternative splicing; Named isoforms=2;
CC
CC
        Name=1; Synonyms=Brain;
CC
          IsoId=070622-1; Sequence=Displayed;
CC
        Name=2; Synonyms=Muscle;
CC
          IsoId=070622-2; Sequence=VSP 005650, VSP 005651;
CC
     -!- TISSUE SPECIFICITY: EXPRESSED PREDOMINANTLY IN NEURAL AND MUSCULAR
CC
        TISSUES.
     -!- SIMILARITY: Contains 1 reticulon domain.
CC
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CC
     EMBL; AF038537; AAC14906.1; -.
DR
    EMBL; AF038537; AAC14907.1; -.
    EMBL; AF038538; AAC14908.1; -.
DR
DR
    EMBL; AF038539; AAC14909.1; -.
DR
    EMBL; AF093624; AAD13195.1; -.
    MGD; MGI:107612; Rtn2.
DR
DR
    InterPro; IPR003388; Reticulon.
DR
    Pfam; PF02453; Reticulon; 1.
DR
    PROSITE; PS50845; RETICULON; 1.
KW
    Endoplasmic reticulum; Alternative splicing; Transmembrane.
FT
    TRANSMEM
                295
                      315
                               POTENTIAL.
FT
    DOMAIN
                272
                      471
                                RETICULON.
FT5
    VARSPLIC
                1
                      267
                                Missing (in isoform 2).
```

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FT
                                 /FTId=VSP 005650.
FT
    VARSPLIC
                268
                       271
                                 PLLL -> MGSK (in isoform 2).
PТ
                                 /FTId=VSP 005651.
     SEQUENCE
               471 AA; 51346 MW; 9BBD8F372CF63AD3 CRC64;
SQ
  Query Match
                         27.9%; Score 197; DB 1; Length 471;
  Best Local Similarity
                        44.3%; Pred. No. 7.2e-12;
          39; Conservative 20; Mismatches
                                                 29; Indels
                                                                0; Gaps
                                                                            0;
          49 VVDLLYWRDIKKTGVVFGASLFLLLSLTVFSIVSVTAYIALALLSVTISFRIYKGVIQAI 108
Qу
              Db
         272 VADLLYWKDTRTSGAVFTGLMASLLCLLHFSIVSVAAHLALLGLCATISLRVYRKVLOAV 331
         109 AKSDEGHPFRAYLESEVAISEELVQKYS 136
QУ
               : | : | | : | | : : : | : : |
         332 HRGDGTNPFQAYLDMDLTLTREQTERLS 359
RESULT 10
T2RD MOUSE
                                  PRT;
TD
     T2RD MOUSE
                   STANDARD;
                                         243 AA.9
AC
     Q9JKA2;
DT
     15-SEP-2003 (Rel. 42, Created)
     15-SEP-2003 (Rel. 42, Last sequence update)
DT
     15-SEP-2003 (Rel. 42, Last annotation update)
DТ
     Taste receptor type 2 member 13 (T2R13) (Taste receptor family B
DE
DE
     member 3) (TRB3) (Fragment).
GN
    TAS2R13.
OS
    Mus musculus (Mouse).
OC
     Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi;
OC
     Mammalia; Eutheria; Rodentia; Sciuroqnathi; Muridae; Murinae; Mus.
OX
     NCBI TaxID=10090;
RN
     [1]
RΡ
     SEQUENCE FROM N.A.
RC
     STRAIN=DBA/2J;
RX
     MEDLINE=20227309; PubMed=10766242;
RA
     Matsunami H., Montmayeur J.-P., Buck L.B.;
     "A family of candidate taste receptors in human and mouse.";
RT.
    Nature 404:601-604(2000).
RN
     [2]
RΡ
     REVIEW.
    MEDLINE=22135574; PubMed=12139982;
RX
RA
    Montmayeur J.-P., Matsunami H.;
RT
     "Receptors for bitter and sweet taste.";
RL
     Curr. Opin. Neurobiol. 12:366-371(2002).
RN
                                                      CRC64
RP
     REVIEW.
RX
    MEDLINE=21634924; PubMed=11696554;
RΑ
    Margolskee R.F.;
RT
     "Molecular mechanisms of bitter and sweet taste transduction.";
RL
    J. Biol. Chem. 277:1-4(2002).
RN
    [4]
RP
    REVIEW.
RX
    MEDLINE=22469025; PubMed=12581520;
RA
     Zhang Y., Hoon M.A., Chandrashekar J., Mueller K.L., Cook B., Wu D.,
RA
     Zuker C.S., Ryba N.J.;
RT
     "Coding of sweet, bitter, and umami tastes: different receptor cells
```

```
RT
    sharing similar signaling pathways.";
RL
    Cell 112:293-301(2003).
CC
    -!- FUNCTION: Receptor that may play a role in the perception of
        bitterness and is gustducin-linked. May play a role in sensing the
CC
CC
        chemical composition of the gastrointestinal content. The activity
CC
        of this receptor may stimulate alpha gustducin, mediate PLC-beta-2
CC
        activation and lead to the gating of TRPM5.
CC
    -!- SUBCELLULAR LOCATION: Integral membrane protein.
CC
    -!- TISSUE SPECIFICITY: Expressed in subsets of taste receptor cells
CC
        of the tongue and palate epithelium and exclusively in gustducin-
CC
        positive cells.
CC
    -!- MISCELLANEOUS: Most taste cells may be activated by a limited
CC
        number of bitter compounds; individual taste cells can
CC
        discriminate among bitter stimuli.
CC
    -!- SIMILARITY: Belongs to family T2R of G-protein coupled receptors.
CC
    -----
CC
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    between the Swiss Institute of Bioinformatics and the EMBL outstation -
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    the European Bioinformatics Institute. There are no restrictions on its
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    entities requires a license agreement (See http://www.isb-sib.ch/announce/
CC
    or send an email to license@isb-sib.ch).
CC
    _____
DR
    EMBL; AF247733; AAF64510.1; -.
    MGD; MGI:1890148; Tas2r13.
DR
DR
    Pfam; PF05296; TAS2R; 1.
KW
    Receptor; G-protein coupled receptor; Transmembrane.
FT
    NON TER
                      1
                1
FT
    DOMAIN
                <1
                      12
                              CYTOPLASMIC (POTENTIAL).
FT
    TRANSMEM
                13
                     33
                              2 (POTENTIAL).
FT
    DOMAIN
               34
                     54
                              EXTRACELLULAR (POTENTIAL).
FT
                     75
    TRANSMEM
               55
                              3 (POTENTIAL).
FT
    DOMAIN
               76
                     99
                              CYTOPLASMIC (POTENTIAL).
FT
    TRANSMEM
              100 120
                              4 (POTENTIAL).
FT
    DOMAIN
              121 150
                             EXTRACELLULAR (POTENTIAL).
FΤ
    TRANSMEM
              151 171
                             5 (POTENTIAL).
FT
    DOMAIN
              172
                    195
                             CYTOPLASMIC (POTENTIAL).
FT
    TRANSMEM
               196 216
                              6 (POTENTIAL).
FT
               217
                    222
                              EXTRACELLULAR (POTENTIAL).
    DOMAIN
FТ
    TRANSMEM
               223 >243
                              7 (POTENTIAL).
FT
              128
    CARBOHYD
                    128
                              N-LINKED (GLCNAC. . .) (POTENTIAL).
FT
    NON TER
               243
                     243
SQ
    SEQUENCE
              243 AA; 28110 MW; D8AD14AF95B9E0B2 CRC64;
 Query Match
                       11.0%; Score 77.5; DB 1; Length 243;
 Best Local Si
                       27.1%; Pred. No. 1.8;
 Matches 32; Conservative 18; Mismatches 47; Indels
                                                         21; Gaps
          17 VYSVSVGMHNLLLLEGRSWQEMDGQKKH---WKDKVVDLLYWRDIKKTGVVFGASLFLLL 73
Qу
            :|| : :|:: || :: || ::|: |
Db
          37 LYSALMTTRKVLIIFNNSWTVIN----HFNIWLATCLSIFYFLKIAN----FSNSIFLSL 88
          74 SLTVFSIVSVTAYIALALLSV-----TISFRIYKGVIQAIAKSDEG-HPFRAYL 121
Qу
               Db
          89 RWRVKTVVSVTLMMSLLLLFVNVLVINTFIVISVDVYKVNTSYSSHSDNNLHISRIFL 146
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RESULT 11
PHSC_ECOLI
                   STANDARD;
                                  PRT:
                                         261 AA.
     PHSC ECOLI
ID
AC
     P77409;
     01-NOV-1997 (Rel. 35, Created)
DT
     01-NOV-1997 (Rel. 35, Last sequence update)
DT
     16-OCT-2001 (Rel. 40, Last annotation update)
DT
DE
     PhsC protein homolog.
    YDHU OR B1670.
GN
OS
    Escherichia coli.
     Bacteria; Proteobacteria; Gammaproteobacteria; Enterobacteriales;
OC
OC
     Enterobacteriaceae; Escherichia.
OX
     NCBI TaxID=562;
RN
     [1]
     SEQUENCE FROM N.A.
RΡ
     STRAIN=K12 / MG1655;
RC
RX
     MEDLINE=97426617; PubMed=9278503;
     Blattner F.R., Plunkett G. III, Bloch C.A., Perna N.T., Burland V.,
RA
     Riley M., Collado-Vides J., Glasner J.D., Rode C.K., Mayhew G.F.,
RA
     Gregor J., Davis N.W., Kirkpatrick H.A., Goeden M.A., Rose D.J.,
RA
RA
     Mau B., Shao Y.;
     "The complete genome sequence of Escherichia coli K-12.";
RT
RL
     Science 277:1453-1474(1997).
RT
     01B
RP
     SEQUENCE FROM N.A.
RC
     STRAIN=K12 / MG1655;
     MEDLINE=97175536; PubMed=9023191;
RX
RA
     Hensel M., Shea J.E., Baeumler A.J., Gleeson C., Blattner F.R.,
RA
     Holden D.W.;
RT
     "Analysis of the boundaries of Salmonella pathogenicity island 2 and
RT
     the corresponding chromosomal region of Escherichia coli K-12.";
RL
     J. Bacteriol. 179:1105-1111(1997).
RN
     [3]
RΡ
     SEQUENCE FROM N.A.
RC
     STRAIN=K12;
     MEDLINE=97251357; PubMed=9097039;
RX
RA
     Aiba H., Baba T., Fujita K., Hayashi K., Inada T., Isono K.,
RA
     Itoh T., Kasai H., Kashimoto K., Kimura S., Kitakawa M.,
RΑ
     Kitagawa M., Makino K., Miki T., Mizobuchi K., Mori H., Mori T.,
RA
     Motomura K., Nakade S., Nakamura Y., Nashimoto H., Nishio Y.,
RA
     Oshima T., Saito N., Sampei G., Seki Y., Sivasundaram S.,
RA
     Tagami H., Takeda J., Takemoto K., Takeuchi Y., Wada C.,
RΑ
     Yamamoto Y., Horiuchi T.;
     "A 570-kb DNA sequence of the Escherichia coli K-12 genome
RT
RT
     corresponding to the 28.0-40.1 min region on the linkage map.";
RL
     DNA Res. 3:363-377(1996).
CC
     -!- SUBCELLULAR LOCATION: Integral membrane protein. Inner membrane
CC
         (Potential).
     -!- SIMILARITY: TO S.TYPHIMURIUM PHSC.
CC
CC
     _____
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    _______
CC
    EMBL; AE000262; AAC74740.1; -.
DR
DR
    EMBL; U68703; AAB47946.1; -.
DR
    EMBL; D90810; BAA15442.1; -.
DR
    PIR; F64924; F64924.
DR
    EcoGene; EG13955; ydhU.
    InterPro; IPR000516; Ni hydr CytB.
DR
DR
    Pfam; PF01292; Ni hydr CYTB; 1.
    Transmembrane; Inner membrane; Complete proteome.
KW
    TRANSMEM
FT
             25 45 POTENTIAL.
                     101
FT
    TRANSMEM
               81
                              POTENTIAL.
FT
    TRANSMEM
             108 128
                              POTENTIAL.
FT
             182 202
                              POTENTIAL.
    TRANSMEM
    TRANSMEM 224 244 POTENTIAL.
FT
    SEQUENCE 261 AA; 29583 MW; 65CF1A45691A0AF3 CRC64;
SO
                       11.0%; Score 77.5; DB 1; Length 261;
 Query Match
 Best Local Similarity 27.6%; Pred. No. 1.9;
         29; Conservative 16; Mismatnhes
                                             48¢) Indels
          22 VGMHNLLLLEGRSWQEMD-GQKKHWKDKVVDLLYWRDIKKTGVVFGASLFLLL----SLT 76
Qу
            44 LGLHALLRARGVKKSATDHGEKIYLYSKAVRLWHWSN-----ALLFVLLLASGLIN 94
Db
          77 VFSIVSVTAYIALALLSVTISFRI---YKGVIQAIAKSDEGHPFR 118
QУ
              Db
          95 HFAMVGATAVKSLVAVHEVCGFLLLACWLGFVLINAVGDNGHHYR 139
RESULT 12
T2R8 MOUSE
ID
    T2R8 MOUSE
                  STANDARD;
                               PRT;
                                      246 AA.
AC
    Q9JKA0;
DT
    15-SEP-2003 (Rel. 42, Created)
DT
    15-SEP-2003 (Rel. 42, Last sequence update)
    15-SEP-2003 (Rel. 42, Last annotation update)
DT
    Taste receptor type 2 member 8 (T2R8) (Taste receptor family B member
DE
    5) (TRB5) (Fragment).
GN
    TAS2R8.
OS
    Mus musculus (Mouse).
    Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi;
OC
    Mamma a; Eutheria; Rodentia; Sciurognathî; lMuridae; Murinae; Mus.
OX
    NCBI TaxID=10090;
RN
    [1]
RP
    SEOUENCE FROM N.A.
RC
    STRAIN=C57BL/6J;
    MEDLINE=20227309; PubMed=10766242;
RX
    Matsunami H., Montmayeur J.-P., Buck L.B.;
    "A family of candidate taste receptors in human and mouse.";
RT
    Nature 404:601-604(2000).
RL
RN
    [2]
RP
    REVIEW.
    MEDLINE=22135574; PubMed=12139982;
RX
RA
    Montmayeur J.-P., Matsunami H.;
    "Receptors for bitter and sweet taste.";
RT
    Curr. Opin. Neurobiol. 12:366-371(2002).
```

```
RN
     [3]
RP
    REVIEW.
    MEDLINE=21634924; PubMed=11696554;
RX
RA
    Margolskee R.F.;
    "Molecular mechanisms of bitter and sweet taste transduction.";
    J. Biol. Chem. 277:1-4(2002).
RI.
RN
     [4]
RP
    REVIEW.
RX
    MEDLINE=22469025; PubMed=12581520;
    Zhang Y., Hoon M.A., Chandrashekar J., Mueller K.L., Cook B., Wu D.,
RA
RA
    Zuker C.S., Ryba N.J.;
    "Coding of sweet, bitter, and umami tastes: different receptor cells
RT
RT
     sharing similar signaling pathways.";
RL
     Cell 112:293-301(2003).
CC
     -!- FUNCTION: Receptor that may play a role in the perception of
        bitterness and is qustducin-linked. May play a role in sensing the
CC
        chemical composition of the gastrointestinal content. The activity
CC
        of this receptor may stimulate alpha gustducin, mediate PLC-beta-2
CC
        activation and lead to the gating of TRPM5.
CC
                    LOCATIONLONAtegral membrane protein.
CC
     -!- TISSUE SPECIFICITY: Expressed in subsets of taste receptor cells
CC
        of the tongue and palate epithelium and exclusively in gustducin-
CC
        positive cells. Expressed in 15% taste bud cells in circumvallate
CC
        and foliate papillae but only in 2% in fungiform papillae.
CC
     -!- MISCELLANEOUS: Most taste cells may be activated by a limited
CC
        number of bitter compounds; individual taste cells can
CC
CC
        discriminate among bitter stimuli.
     -!- SIMILARITY: Belongs to family T2R of G-protein coupled receptors.
CC
     ______
CC
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     the European Bioinformatics Institut@KYTThere are no rest
                                                                  ns on its
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     modified and this statement is not removed. Usage by and for commercial
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     or send an email to license@isb-sib.ch).
CC
     ______
CC
DR
     EMBL; AF247735; AAF64512.1; -.
DR
     MGD; MGI:1890259; Tas2r8.
DR
     Pfam; PF05296; TAS2R; 1.
     Receptor; G-protein coupled receptor; Transmembrane.
KW
FT
     NON TER
                  1
                        1
                                CYTOPLASMIC (POTENTIAL).
FT
     DOMAIN
                  1
                        15
FT
     TRANSMEM
                 16
                        36
                                2 (POTENTIAL).
                                EXTRACELLULAR (POTENTIAL).
FT
     DOMAIN
                 37
                        59
FT
                 60
                       80
                                3 (POTENTIAL).
     TRANSMEM
FТ
     DOMAIN
                81
                       102
                                CYTOPLASMIC (POTENTIAL).
                103
                       123
FT
     TRANSMEM
                                 4 (POTENTIAL).
FT
     DOMAINC
                M N.A.459XN
                                EXTRACELLULAR (POTENTIAL).
FT
     TRANSMEM
                154
                       174
                                 5 (POTENTIAL).
                175
                                CYTOPLASMIC (POTENTIAL).
FT
     DOMAIN
                       198
                199
                       219
                                6 (POTENTIAL).
FT
     TRANSMEM
FΤ
     DOMAIN
                220
                       225
                                EXTRACELLULAR (POTENTIAL).
FT
     TRANSMEM
                226
                       246
                                7 (POTENTIAL).
FT
     NON TER
                246
                       246
     SEQUENCE
               246 AA; 28430 MW; 8B8F96F8A62E4474 CRC64;
SO
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10.7%; Score 75.5; DB 1; Length 246;
   Query Match
   Best Local Similarity 27.1%; Pred. No. 2.9;
                                                                                                              21; Gaps
                  32; Conservative 18; Mismatches 47; Indels
                                                                                                                                    5;
                  17 VYSVSVGMHNLLLLEGRSWQEMDGQKKH---WKDKVVDLLYWRDIKKTGVVFGASLFLLL 73
Qу
                        :|| : :|: || :: || :: || :|: |
                  40 LYSALMTTRKVLIIFNNSWTVIN----HFNIWLATCLSIFYFLMIAN----FSNSIFLSL 91
Db
                  74 SLTVFSIVSVTAYIALALLSV-----TISFRIYKGVIQAIAKSDEG-HPFRAYL 121
QУ
                             1 :: | | | | : : | | | | : | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : | | : 
                  92 RWRVKTVVSVTLLMSLLLLFVNVLVINTFIVISVDVYKVNTSYSSHSDNNIHISRIFL 149
Dh
RESULT 13
G6PI HELPY
        G6PI HELPY
                                 STANDARD;
                                                     PRT;
                                                                        545 AA.
ID
AC
        025781;
        15-JUL-1998 (Rel. 36, Created)
DT
        15-JUL-1998 (Rel. 36, Last sequence update)
DT
                                                   1,afimeLatdohXOpdate)
DT
        28-FEB-2003 (Rel.
        Glucose-6-phosphate isomerase (EC 5.3.1.9) (GPI) (Phosphoglucose
DE
        isomerase) (PGI) (Phosphohexose isomerase) (PHI).
DE
        PGI OR HP1166.
GN
OS
        Helicobacter pylori (Campylobacter pylori).
        Bacteria; Proteobacteria; Epsilonproteobacteria; Campylobacterales;
OC
        Helicobacteraceae; Helicobacter.
OC
OX
        NCBI TaxID=210;
RN
         [1]
RP
        SEQUENCE FROM N.A.
        STRAIN=26695 / ATCC 700392;
RC
RX
        MEDLINE=97394467; PubMed=9252185;
RA
        Tomb J.-F., White O., Kerlavage A.R., Clayton R.A., Sutton G.G.,
RA
         Fleischmann R.D., Ketchum K.A., Klenk H.-P., Gill S., Dougherty B.A.,
RA
        Nelson K., Quackenbush J., Zhou L., Kirkness E.F., Peterson S.,
RA
        Loftus B., Richardson D., Dodson R., Khalak H.G., Glodek A.,
        McKenney K., FitzGerald L.M., Lee N., Adams M.D., Hickey E.K.,
RA
        Berg D.E., Gocayne J.D., Utterback T.R., Peterson J.D., Kelley J.M.,
RA
         Cotton M.D., Weidman J.M.L.; 00XB C., Bowman C., Watthey L., Wallin E.,
RA
RA
        Hayes W.S., Borodovsky M., Karp P.D., Smith H.O., Fraser C.M.,
RA
         Venter J.C.;
RT
        "The complete genome sequence of the gastric pathogen Helicobacter
RT
        pylori.";
RL
        Nature 388:539-547(1997).
CC
         -!- CATALYTIC ACTIVITY: D-qlucose 6-phosphate = D-fructose 6-
CC
               phosphate.
CC
         -!- PATHWAY: Involved in glycolysis and in gluconeogenesis.
CC
         -!- SUBCELLULAR LOCATION: Cytoplasmic (By similarity).
CC
         -!- SIMILARITY: BELONGS TO THE GPI FAMILY.
         CC
CC
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         use by non-profit institutions as long a Cts c6Atent is in no way
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         or send an email to license@isb-sib.ch).
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EMBL; AE000622; AAD08211.1; -.
DR
DR
    PIR; F64665; F64665.
    HSSP; Q9N1E2; 1HOX.
DR
    TIGR; HP1166; -.
DR
    HAMAP; MF 00473; -; 1.
DR
    InterPro; IPR001672; G6P Isomerase.
DR
DR
    Pfam; PF00342; PGI; 1.
    PRINTS; PR00662; G6PISOMERASE.
DR
    PROSITE; PS00765; P GLUCOSE ISOMERASE 1; 1.
DR
    PROSITE; PS00174; P GLUCOSE ISOMERASE_2; 1.
DR
    Isomerase; Gluconeogenesis; Glycolysis; Complete proteome.
KW
                                BY SIMILARITY.
                     382
FT
    ACT SITE
               382
                510
                       510
                                BY SIMILARITY.
FT
    ACT SITE
SQ
    SEQUENCE
               545 AA; 62487 MW; BDC68D1625190236 CRC64;
                         10.7%; Score 75.5; DB 1; Length 545;
  Query Match
  Best Local Similarity 25.3%; Pred. No. 6.7;
                                               44; Indels
                                                             45; Gaps
  Matches
           37; Conservative 20; Mismatches
          23 GMHNLL-----LLEGRSWQEMDGQ---KKHWKDKVVDLLYWRDIKKTGVVFGASL 69
Qу
             | | :|
                             ::|:|::|
                                                411 GHHEILFSNVLAQAQAFMKGKSYEEALGELLFKGLDKDEAKDLAHHR-----VFFGNRP 464
Db
          70 FLLLSLTVFSIVSVTAYIALALLSVTISFRIYKGVIQAIAKSD----- 112
QУ
               465 SNILLLEKISPSNIGALVALYEHKVFV-----QGVIWDINSFDQWGVELGKELAVPILQE 519
Db
         113 -EGHPFRAYLESEVAISEELVQKYSN 137
Οv
              :: |:: | |
         520 LEGHKSNAYFDSS---TKHLIELYKN 542
Db
RESULT 14
G6PI HELPJ
     G6PI HELPJ
                   STANDARD;
                                  PRT;
                                        545 AA.
AC
     Q9ZK49;
     16-OCT-2001 (Rel. 40, Created)
DT
     16-OCT-2001 (Rel. 40, Last sequence update)
DT
     28-FEB-2003 (Rel. 41, Last annotation update)
DT
     Glucose-6-phosphate isomerase (EC 5.3.1.9) (GPI) (Phosphoglucose
DΕ
DΕ
     isomerase) (PGI) (Phosphohexose isomerase) (PHI).
GN
     PGI OR JHP1093.
OS
     Helicobacter pylori J99 (Campylobacter pylori J99).
OC
     Bacteria; Proteobacteria; Epsilonproteobacteria; Campylobacterales;
     Helicobacteraceae; Helicobacter.
OC
OX
     NCBI TaxID=85963;
RN
     [1]
RP
     SEQUENCE FROM N.A.
     MEDLINE=99120557; PubMed=9923682;
RX
     Alm R.A., Ling L.-S.L., Moir D.T., King B.L., Brown E.D., Doig P.C.,
RΑ
RA
     Smith D.R., Noonan B., Guild B.C., deJonge B.L., Carmel G.,
RA
     Tummino P.J., Caruso A., Uria-Nickelsen M., Mills D.M., Ives C.,
RA
     Gibson R., Merberg D., Mills S.D., Jiang Q., Taylor D.E., Vovis G.F.,
RA
     Trust T.J.;
     "Genomic sequence comparison of two unrelated isolates of the human
RT
RT
     gastric pathogen Helicobacter pylori.";
RL
     Nature 397:176-180(1999).
```

```
CC
       phosphate.
    -!- PATHWAY: Involved in glycolysis and in gluconeogenesis.
CC
    -!- SUBCELLULAR LOCATION: Cytoplasmic (By similarity).
CC
    -!- SIMILARITY: BELONGS TO THE GPI FAMILY.
CC
    ______
CC
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    the European Bioinformatics Institute. There are no restrictions on its
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    or send an email to license@isb-sib.ch).
CC
    _____
CC
DR
    EMBL; AE001536; AAD06664.1; -.
DR
    PIR; E71851; E71851.
    HSSP; Q9N1E2; 1HOX.
DR
DR
    HAMAP; MF_00473; -; 1.
    InterPro; IPR001672; G6P Isomerase.
DR
    Pfam; PF00342; PGI; 1.
DR
DR
    PRINTS; PR00662; G6PISOMERASE.
    PROSITE; PS00765; P GLUCOSE ISOMERASE 1; 1.
DR
    PROSITE; PS00174; P GLUCOSE ISOMERASE 2; 1.
DR
    Isomerase; Gluconeogenesis; Glycolysis; Complete proteome.
KW
             382 382
FT
                             BY SIMILARITY.
    ACT SITE
FT
    ACT SITE
              510
                    510
                             BY SIMILARITY.
    SEQUENCE 545 AA; 62302 MW; 7DB544D95FD1D237 CRC64;
SQ
                      10.6%; Score 74.5; DB 1; Length 545;
 Query Match
 Best Local Similarity 25.3%; Pred. No. 8.4;
         37; Conservative 19; Mismatches
 Matches
                                            45; Indels
                                                        45; Gaps
                                                                    6;
         23 GMHNLL-----LLEGRSWQEMDGQ---KKHWKDKVVDLLYWRDIKKTGVVFGASL 69
Qу
            | | :|
                           Db
        411 GHHEILFSNVLAQAQAFMKGKSYEEALGELLSKGLDKDEAKDLAHHR-----VFFGNRP 464
         70 FLLLSLTVFSIVSVTAYIALALLSVTISFRIYKGVIQAIAKSD------ 112
Qу
              465 SNILLLEKISPSNIGALVALYEHKVFV-----QGVIWDINSFDQWGVELGKELAVPILQE 519
Db
        113 -EGHPFRAYLESEVAISEELVOKYSN 137
Qу
                 Db
         520 LEGHKSNAYFDSS---TRHLIELYKN 542
RESULT 15
YC73 HAEIN
TD
    YC73 HAEIN
                 STANDARD;
                              PRT:
                                     268 AA.
AC
    P44150;
DT
    01-NOV-1995 (Rel. 32, Created)
DT
    01-NOV-1995 (Rel. 32, Last sequence update)
DT
    28-FEB-2003 (Rel. 41, Last annotation update)
DE
    Hypothetical protein HI1273.
GN
    HI1273.
OS
    Haemophilus influenzae.
OC
    Bacteria; Proteobacteria; Gammaproteobacteria; Pasteurellales;
OC
    Pasteurellaceae; Haemophilus.
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-!- CATALYTIC ACTIVITY: D-glucose 6-phosphate = D-fructose 6-

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OX
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RN
    [1]
RP
    SEQUENCE FROM N.A.
RC
    STRAIN=Rd / KW20 / ATCC 51907;
RX
    MEDLINE=95350630; PubMed=7542800;
    Fleischmann R.D., Adams M.D., White O., Clayton R.A., Kirkness E.F.,
RA
    Kerlavage A.R., Bult C.J., Tomb J.-F., Dougherty B.A., Merrick J.M.,
RΑ
    McKenney K., Sutton G., Fitzhugh W., Fields C.A., Gocayne J.D.,
RA
    Scott J.D., Shirley R., Liu L.-I., Glodek A., Kelley J.M.,
RA
    Weidman J.F., Phillips C.A., Spriggs T., Hedblom E., Cotton M.D.,
RA
    Utterback T.R., Hanna M.C., Nguyen D.T., Saudek D.M., Brandon R.C.,
RA
RA
    Fine L.D., Fritchman J.L., Fuhrmann J.L., Geoghagen N.S.M.,
    Gnehm C.L., McDonald L.A., Small K.V., Fraser C.M., Smith H.O.,
RA
    Venter J.C.;
RA
     "Whole-genome random sequencing and assembly of Haemophilus influenzae
RT
RT
    Science 269:496-512(1995).
RL
    ______
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CC
     EMBL; U32807; AAC22921.1; -.
DR
    PIR; F64024; F64024.
DR
    TIGR; HI1273; -.
DR
DR
     InterPro; IPR000051; SAM bind.
    Hypothetical protein; Complete proteome.
KW
SQ
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  Query Match
                        10.5%; Score 74; DB 1; Length 268;
  Best Local Similarity
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           29; Conservative 15; Mismatches
                                               49; Indels
                                                            22; Gaps
                                                                        4;
           8 VSCLRENFAVYSVSVGMHNLLLLEGRSWQEMDGQKKHWKDKVVDLLYWRDIKKTGVVFGA 67
Qу
                    | :
                           | : | : |
                                              : ||
          94 LDCL----AQFKQKFGLHHLTTFH------KSWADN-----WDDVPQADVVLAS 132
Db
          68 SLFLLLSL-TVFSIVSVTAYIALALLSVTISFRIYKGVIQAIAKSDEGHPFRAYL 121
Qу
                133 RSTLVDDLDDMIEKLCAKAKKRVFLTSVTQRHFLDEGVFEAIGREDIGFPTYIYL 187
Db
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Search completed: January 22, 2004, 16:32:01 Job time: 4.35178 secs

 $\sim$